



# Caring without Borders

Espnic 2013  
June 12-15, 2013  
Rotterdam, Netherlands



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<input type="checkbox"/>	No, nothing to disclose
<input checked="" type="checkbox"/>	Yes, please specify:

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Maquet	X	X		X				

## Off-Label Product Use

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<input type="checkbox"/>	Yes, please specify:

# ***Physiological Concepts Behind Neurally Adjusted Ventilatory Assist (NAVA) and Non-Invasive NAVA (NIV-NAVA)***

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St. Michael's Hospital, Toronto  
Department of Pediatrics, University of Toronto

- [www.VentQuest.ca](http://www.VentQuest.ca)
- Jennifer.beck@rogers.com

# Neural Control of Artificial Muscles

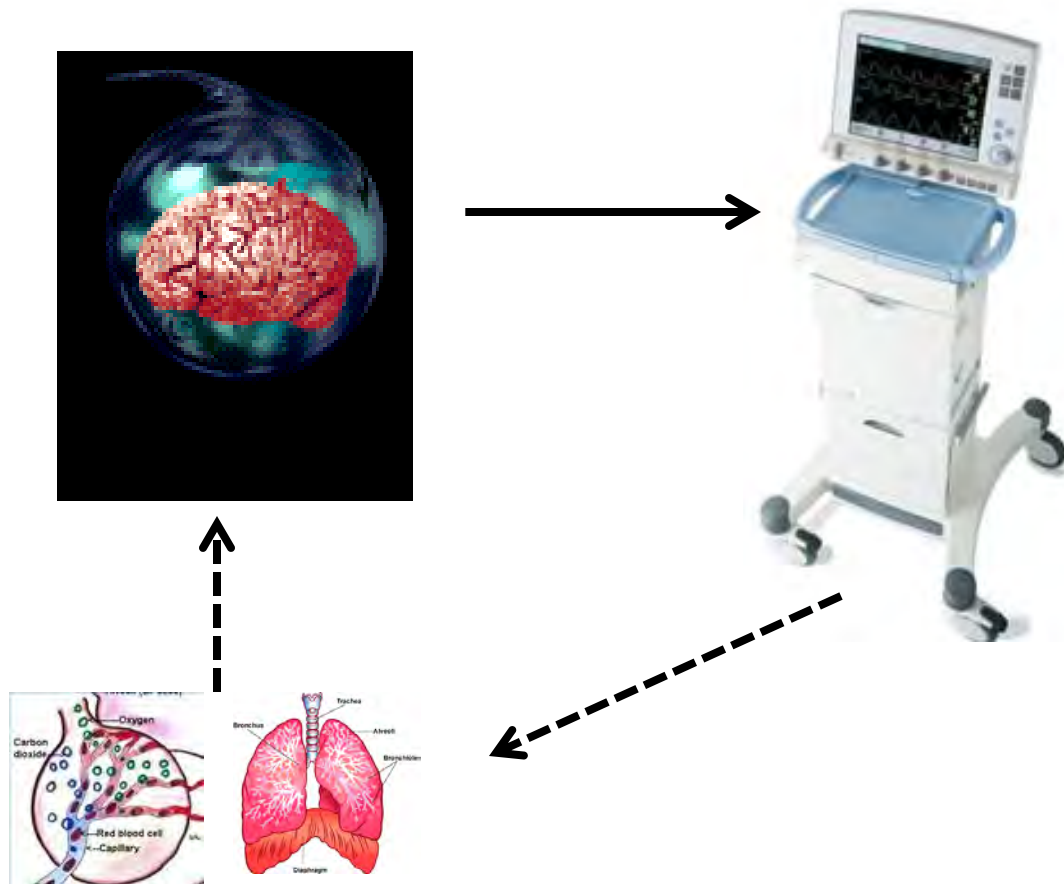


Jan 2010



Time Magazine Jan 2008 (European Ed)

# Neurally Controlled Mechanical Ventilation



# Unique Features about Infants

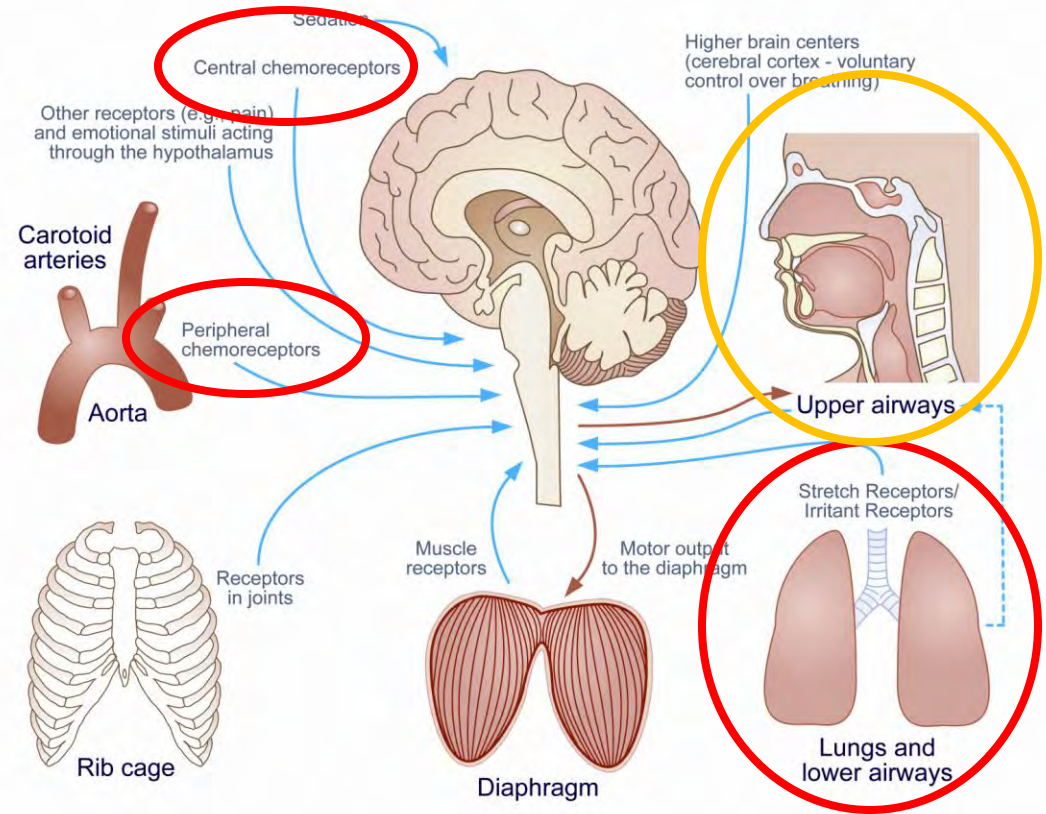
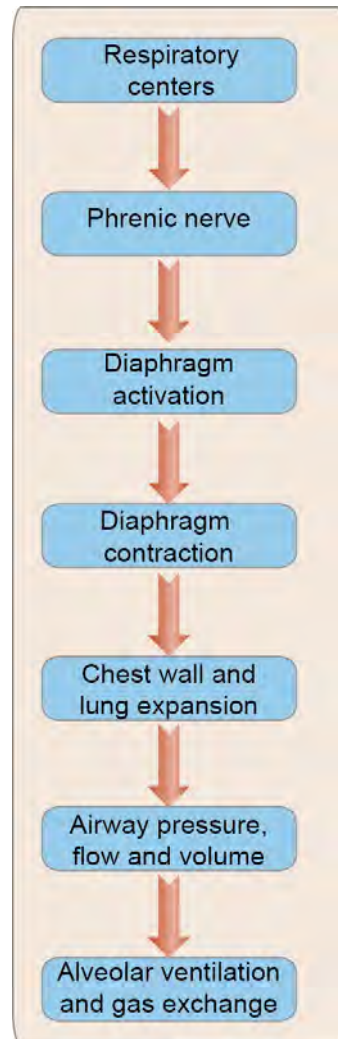
- Surfactant deficiency and compliant chest wall make that the lung has a tendency to de-recruit
- Strong vagal reflexes
- Apnea of prematurity
- Uncuffed ET tubes



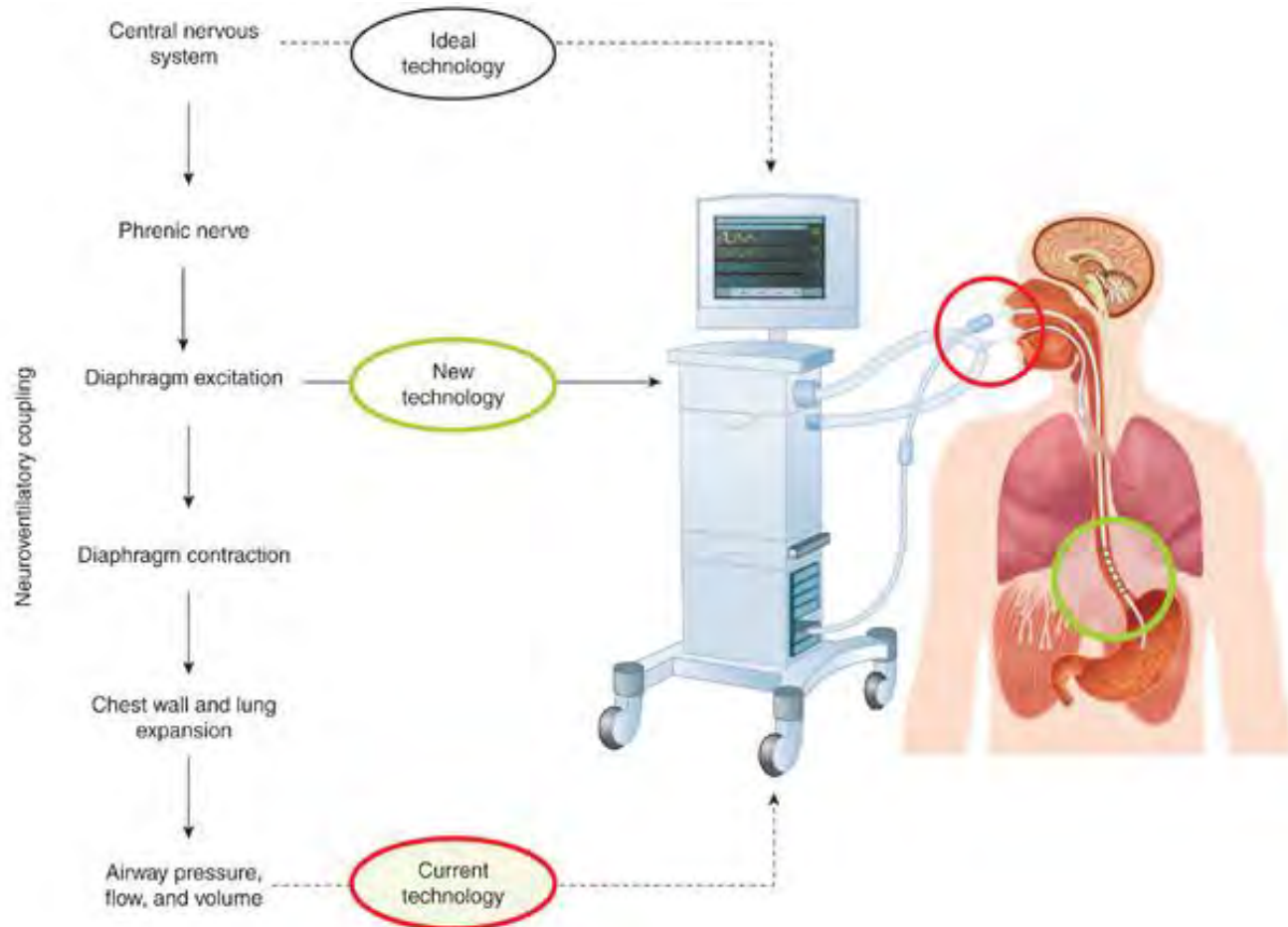
“Maggie” 500 g at birth (with permission from the parents)



# Breathing is Neurally Regulated

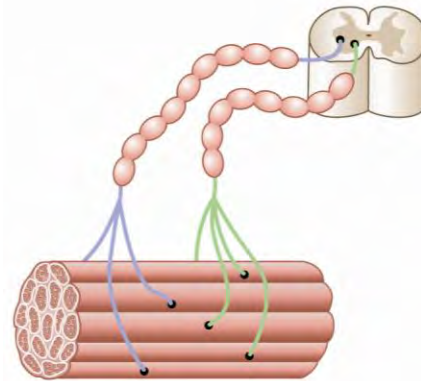
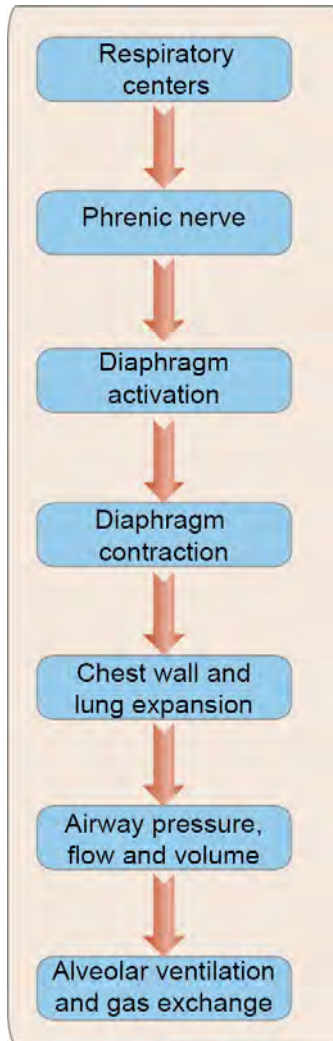


# General Principles of NAVA

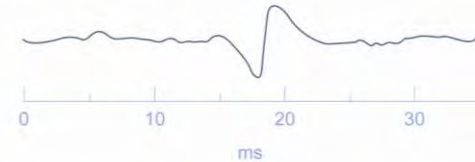




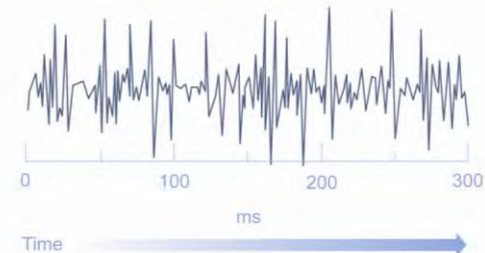
# EMG of Skeletal Muscle



Single motor unit action potential

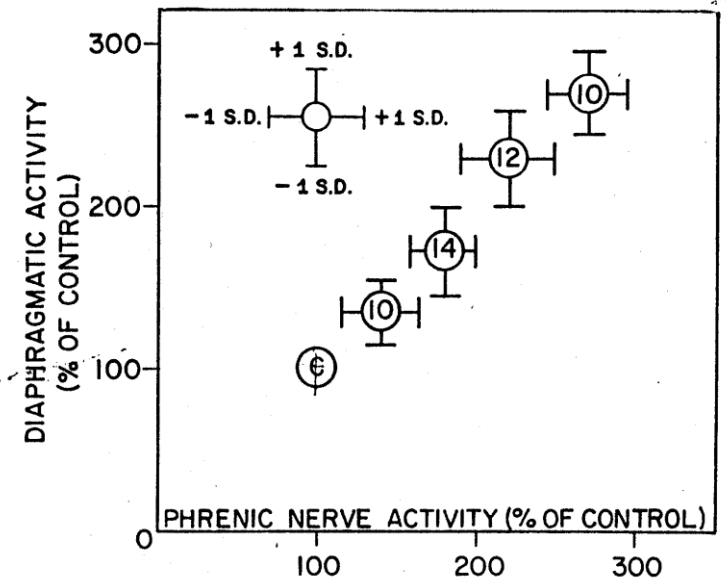
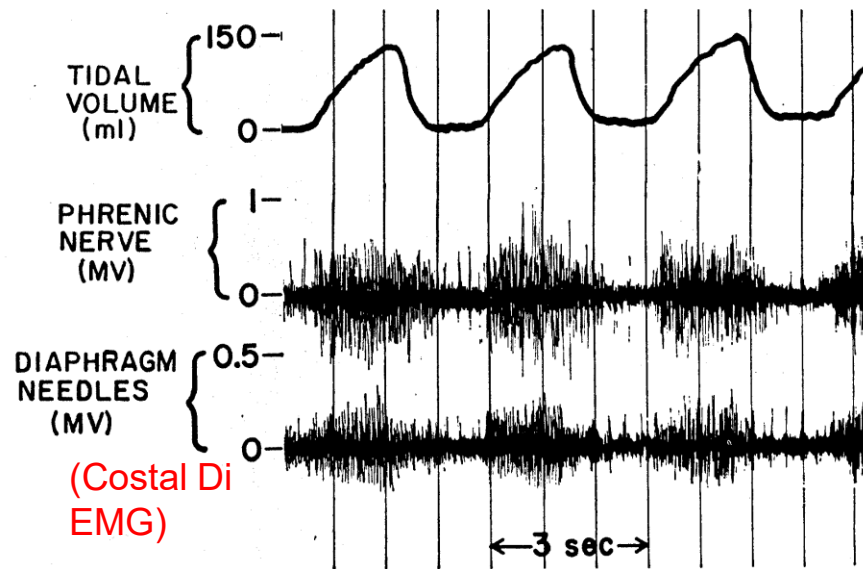


Multiple motor unit interference pattern

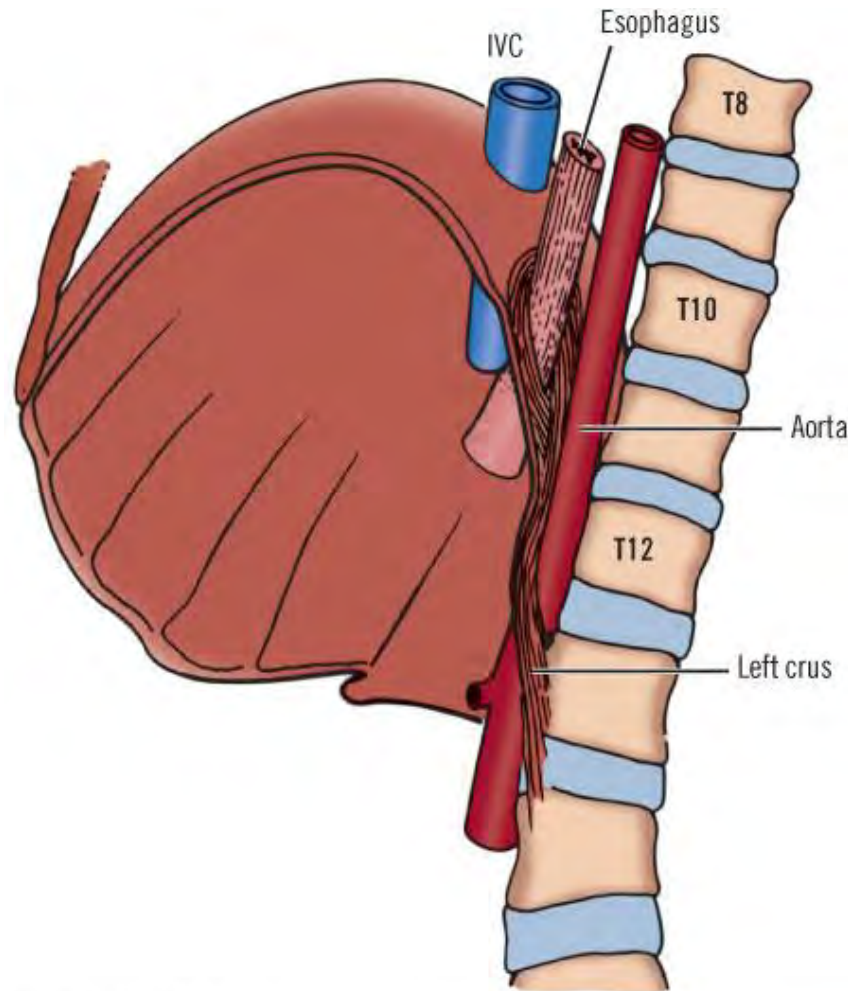


“EMG”

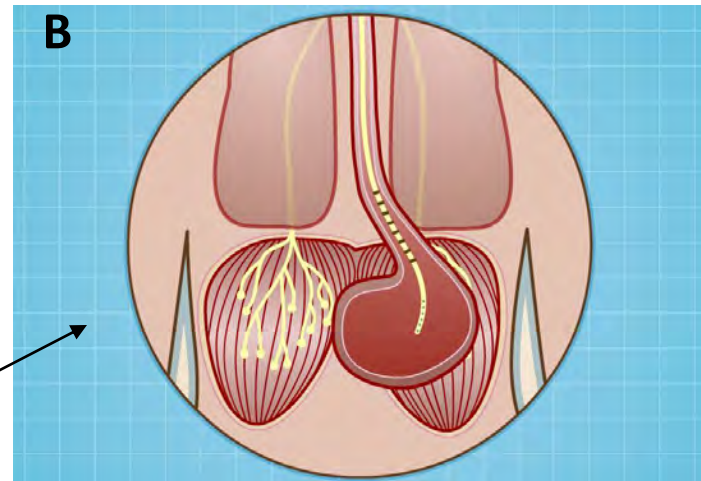
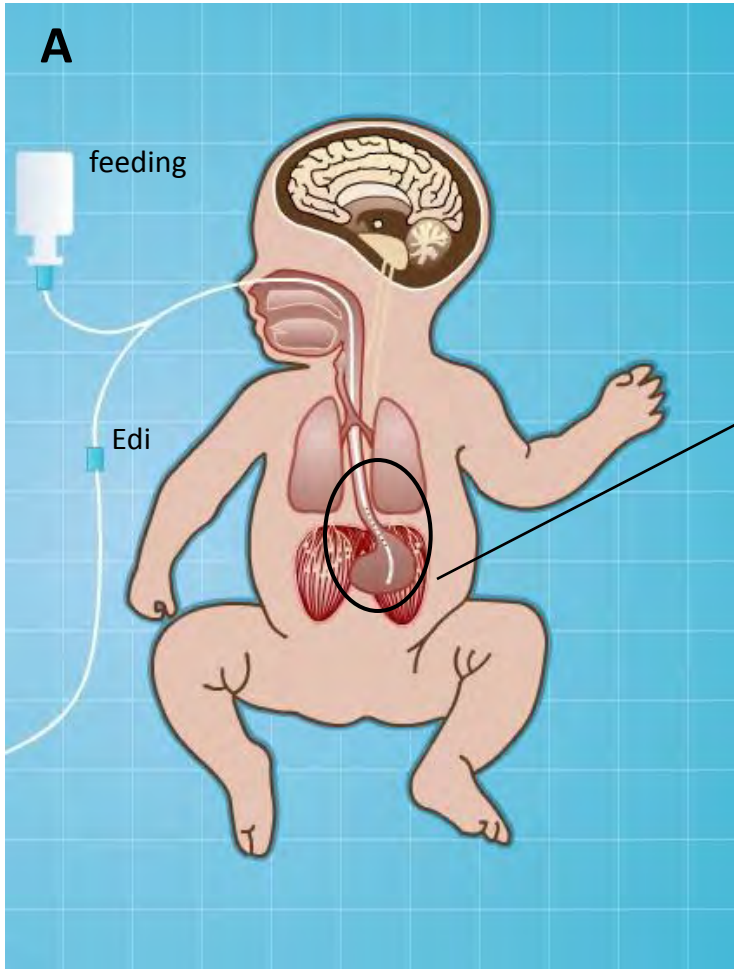
# EMG is the Neural Drive to the Diaphragm



# How to Measure Diaphragm EMG in Humans?



# Modified Feeding Tube



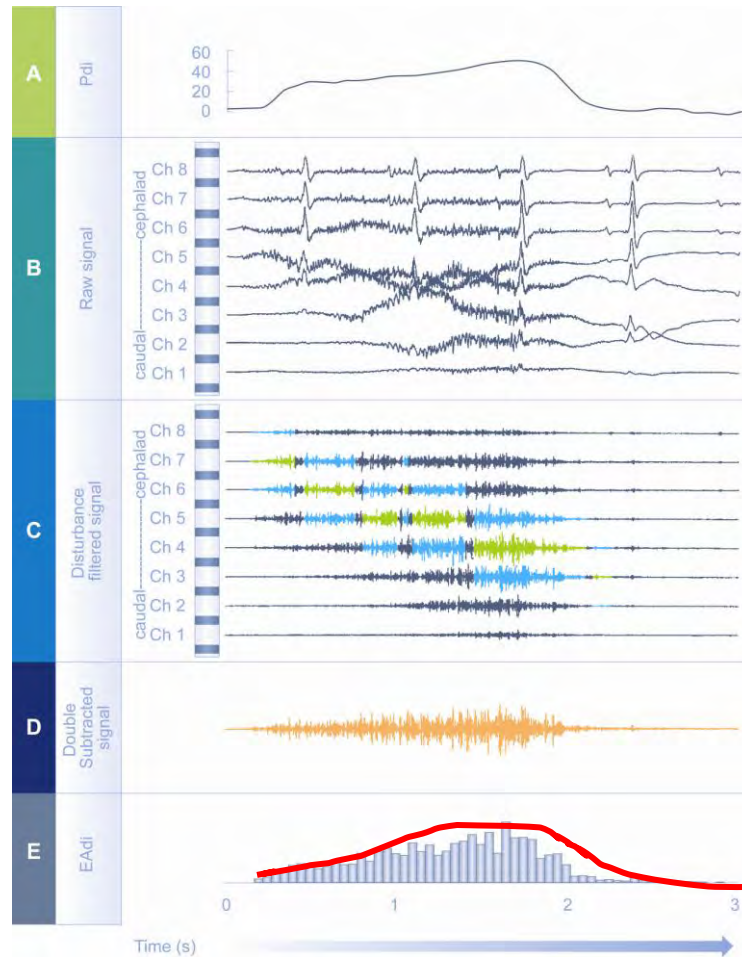
# Electrode Positioning

- NEX
- Calculate prediction
- Insert to prediction
- Positioning window (verification)
- Secure and record final position

## Validation studies:

Barwing, ICM (2009); [Green, Respir Care \(2011\)](#); Barwing, ICM (2011); [Duyndam, Nursing in Crit Care \(2013\)](#); [Stein, J Perinatology \(2013\)](#), [Vignaux, PCCM, In Press \(2013\)](#)

# EMG to Edi Waveform

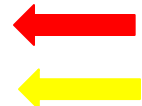
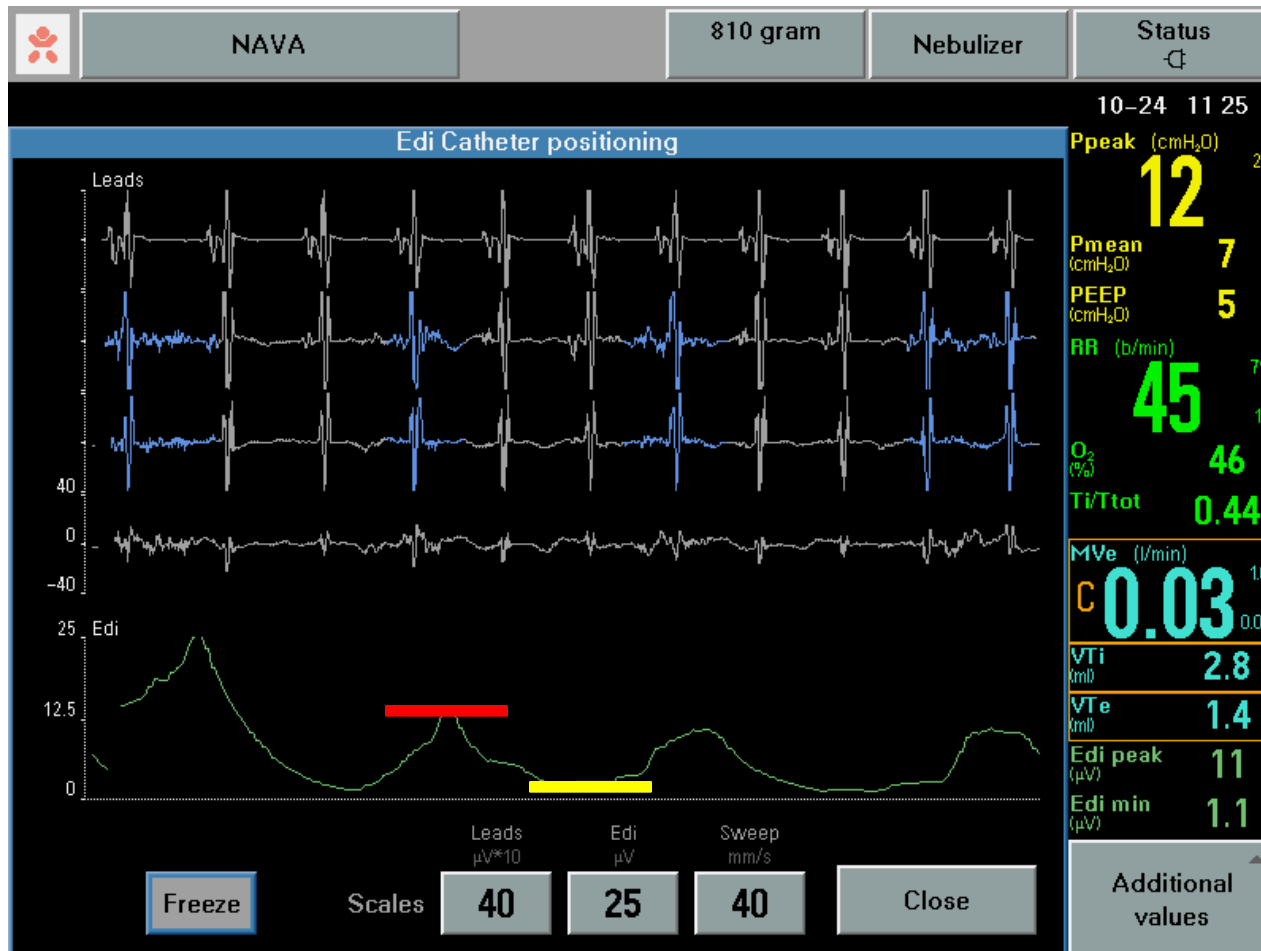




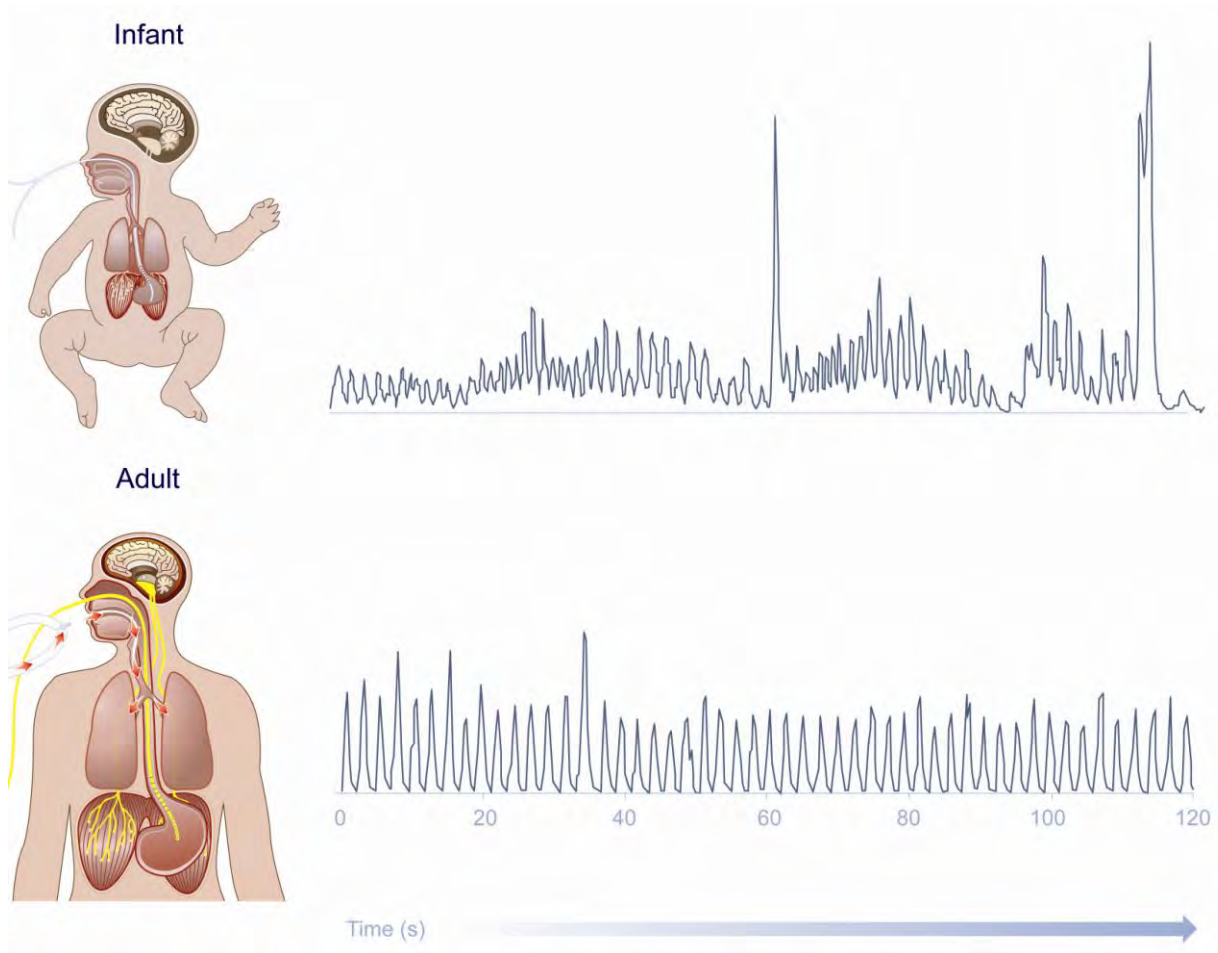
# Edi waveform

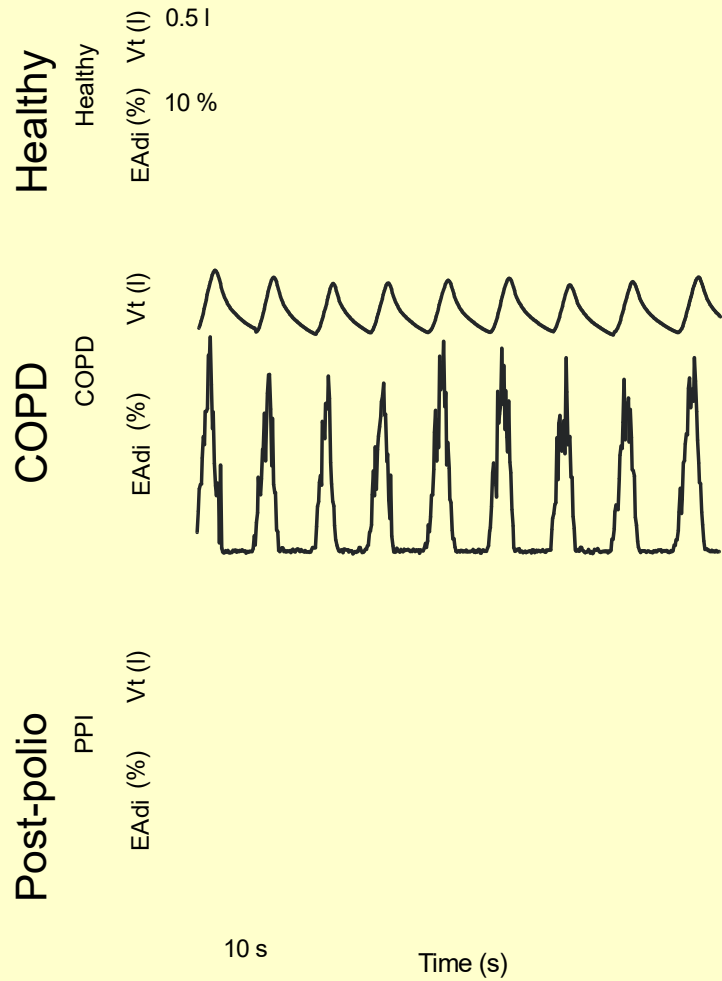
4 leads

Edi



# Edi in Infants and Adult





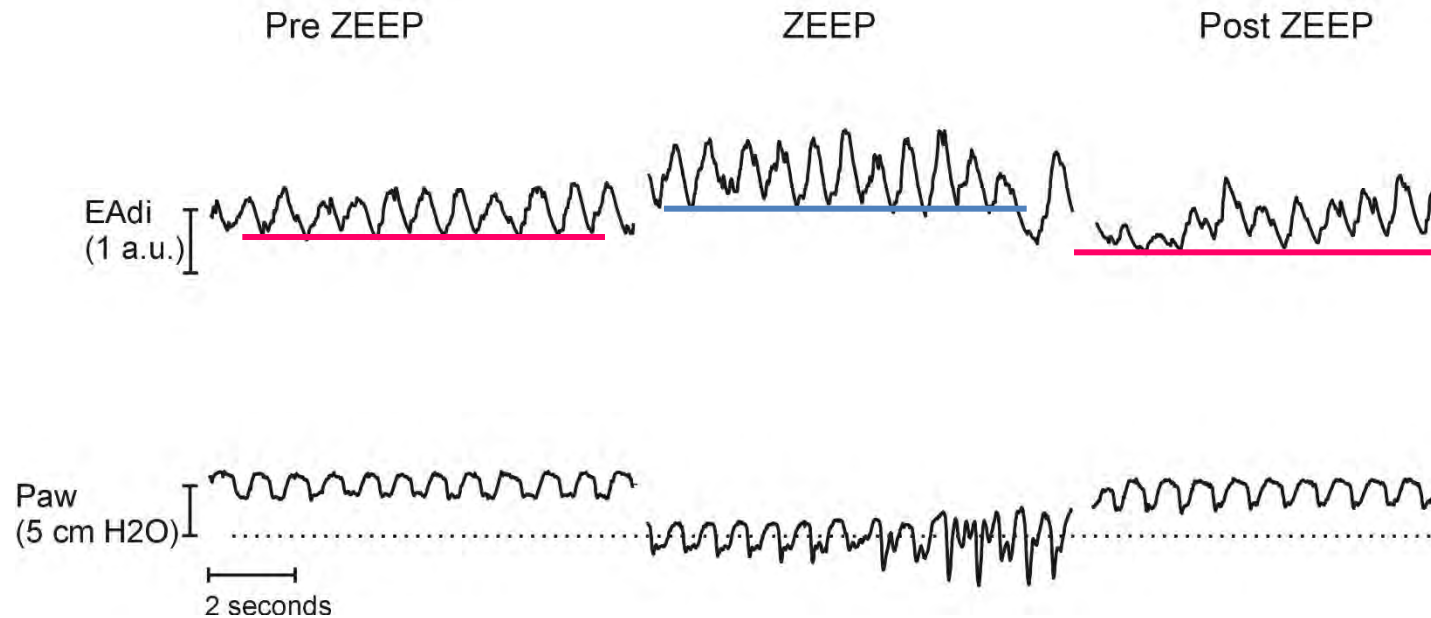
peak

- Reduced level of assist
- Increased Respiratory Load
- Weakness of Diaphragm
- Increased CO<sub>2</sub>
- Reduced sedation

# Published Edi<sub>pk</sub> (on Servoi)

Catheter size	Condition	Mean Edipk (uV)	Lowest mean value Edi pk (uV)	Highest mean value Edi pk (uv)	# studies	# patients
16	Intubated on NAVA	10.4	4	15	16	204
16	NIV-NAVA	19.2	11	33	4	53
8	Intubated on NAVA	9.1	5	11	4	44
8	NIV-NAVA	27	20	34	2	15
6	Intubated on NAVA	9.4	7.4	11.4	3	47
6	NIV-NAVA				0	0
6	No assist	11	10	16	1	3 (healthy)
ALL		13.8	4	34	34	507

# $Edi_{min}$

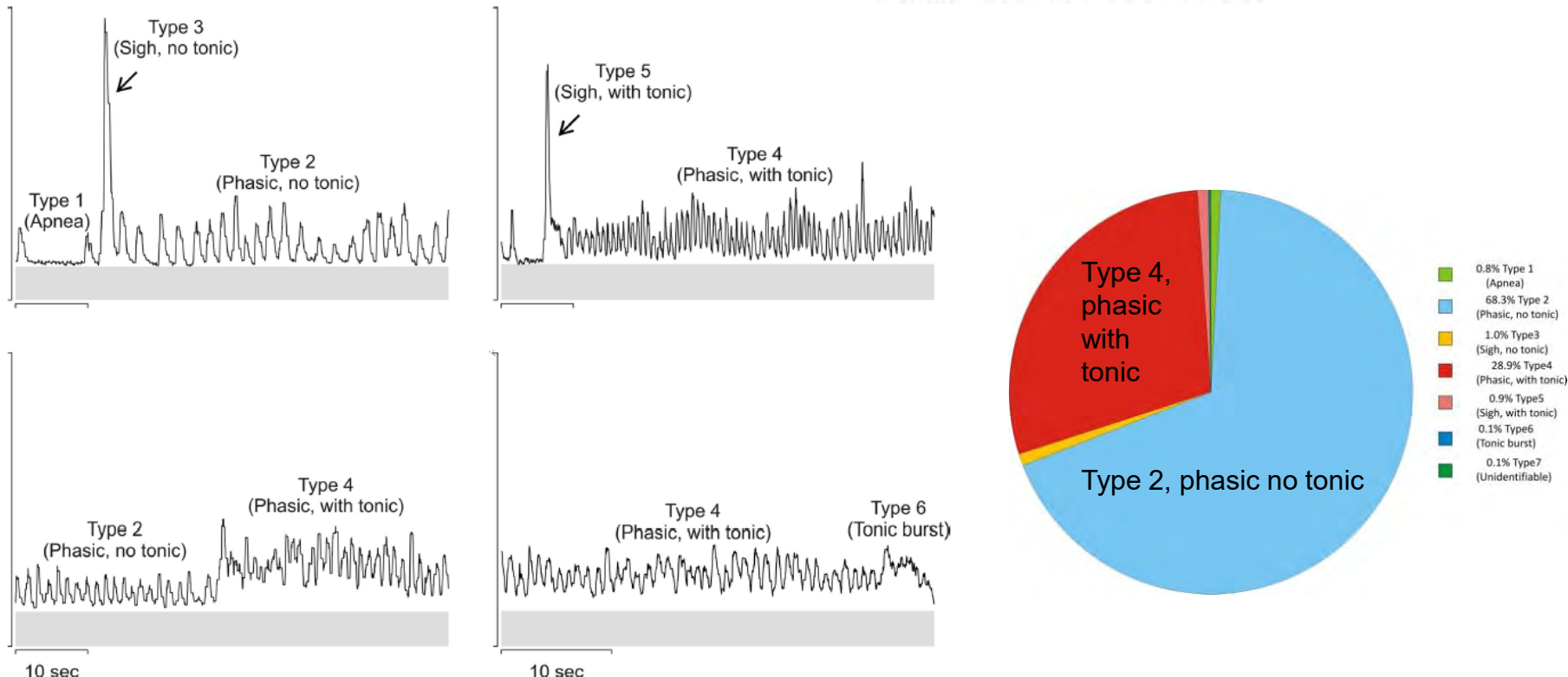


*Emeriaud et al, Ped Res, 2006*

- De-recruitment below FRC
- Liquid/edema in the lung
- ??

## Characterization of Neural Breathing Pattern in Spontaneously Breathing Preterm Infants

JENNIFER BECK, MAUREEN REILLY, GIACOMO GRASSELLI, HAIBO QUI, ARTHUR S. SLUTSKY, MICHAEL S. DUNN,  
 AND CHRISTER A. SINDERBY *Pediatr Res* 70: 607–613, 2011



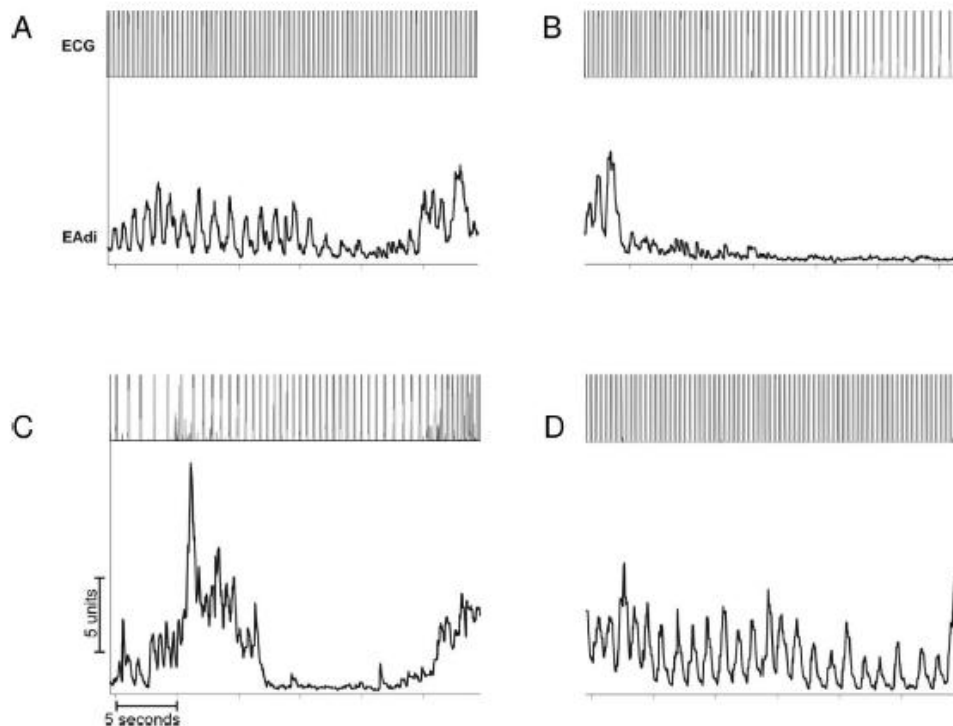
**Figure 4.** Representative tracings of the different neural breathing patterns on the same day: each panel demonstrates the EAdi waveform obtained in one subject on the same recording day, and demonstrates examples of the different breathing pattern types (1–7, see Methods).

N = 10  
 Mean weight = 1400 g  
 Mean age = 8 days  
 GA at birth: 31 wks

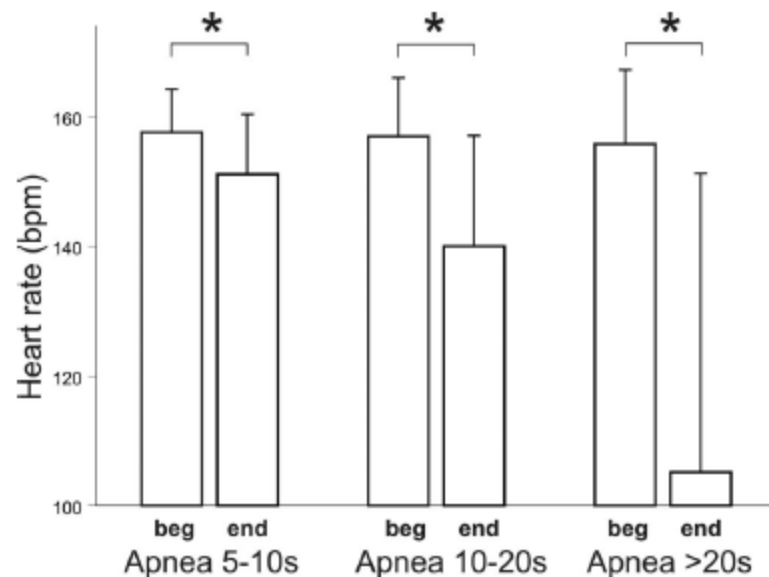


# Characterization of Neural Breathing Pattern in Spontaneously Breathing Preterm Infants

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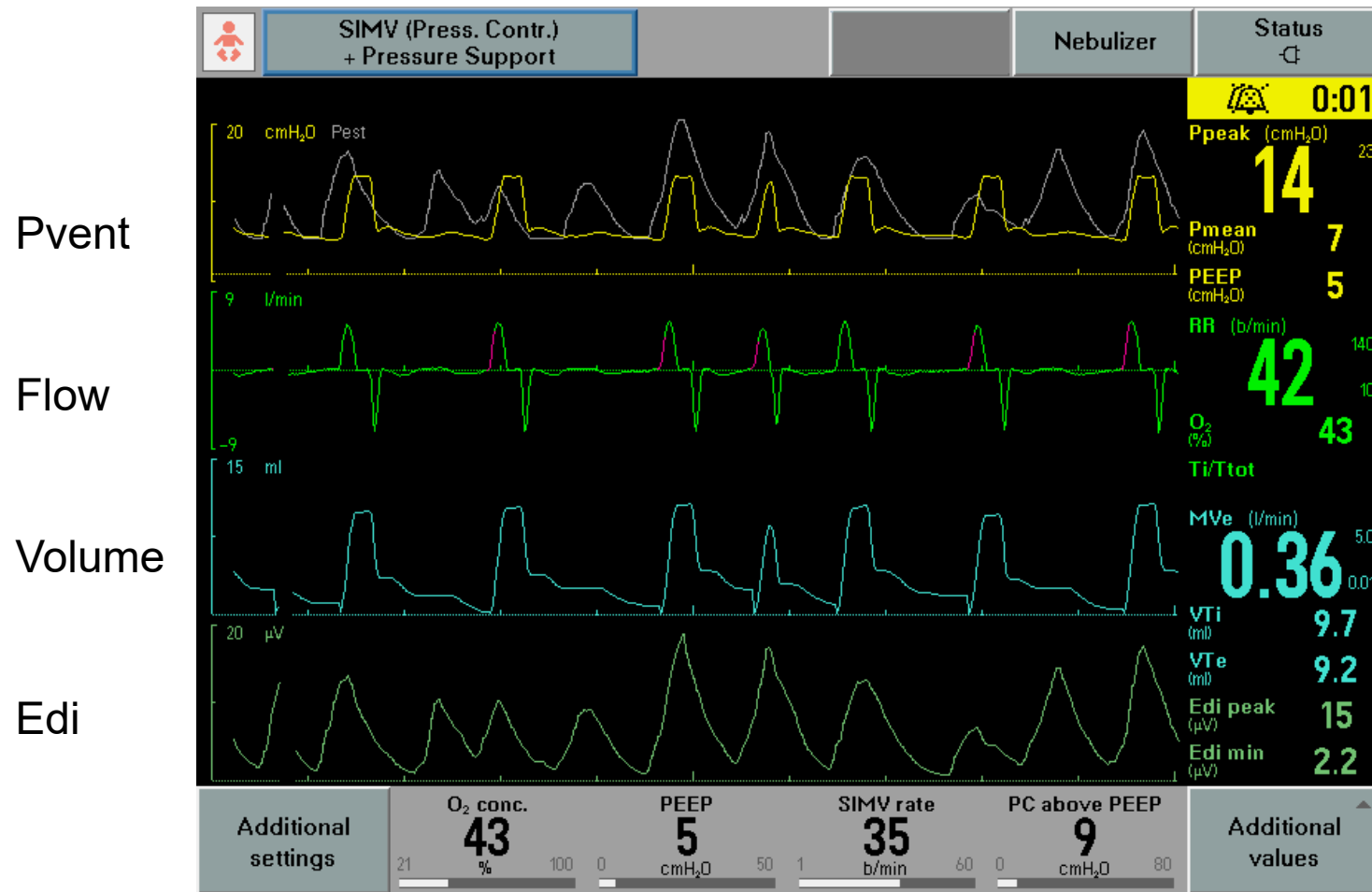
**Figure 6.** Representative tracings of EAdi and ECG in one subject: time sequence is Figure 6A to D. Each panel demonstrates the EAdi waveform (bottom in each panel) and the ECG. (A) = 30 s before the apnea, heart rate was 153, and saturation 98%. (B) = slowing in heart rate at the same time as the central apnea. (C) = Heart rate 66, saturation 82%, and the nurse aroused the baby (D). Three minutes later, HR = heart rate 159 and saturation = 99%.



**Figure 7.** Relationship between central apnea duration and heart rate: group data demonstrating that longer apneas are associated with greater reductions in heart rate. \*Statistical significance from beginning to end of apnea.

N = 10  
Mean weight = 1400 g  
Mean age = 8 days  
GA at birth: 31 wks

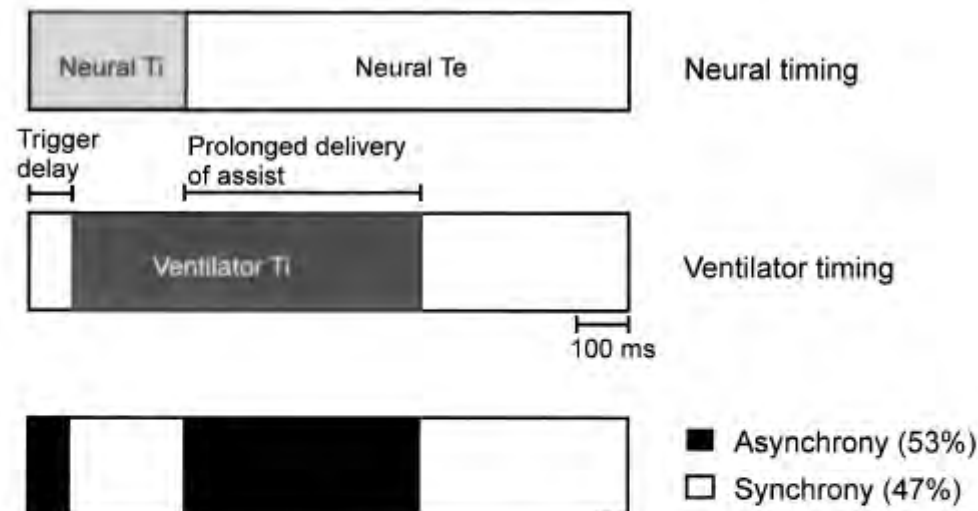
# Edi and Patient-ventilator Interaction



## Prolonged Neural Expiratory Time Induced by Mechanical Ventilation in Infants

JENNIFER BECK, MARISA TUCCI, GUILLAUME EMERIAUD, JACQUES LACROIX, AND  
 CHRISTER SINDERBY

*Pediatric Intensive Care Unit, Department of Pediatrics and Hôpital Sainte-Justine Research Center, Université de Montréal, Montreal, Quebec H3T 1C5 [J.B., M.T., G.E., J.L.]; Department of Newborn and Developmental Pediatrics, Sunnybrook and Women's College Health Sciences Centre, Toronto, Ontario M5S 1B2 [J.B.]; and Department of Critical Care Medicine, St-Michael's Hospital, Toronto, Ontario M5B 1W8 [C.S.], Canada*



N = 14  
 Mean weight = 3.9 kg  
 Mean age = 2.3 mos

**Figure 4.** Patient-ventilator interaction during mandatory breaths. Schematic representation of patient neural timing (*upper bar*) and ventilator timing (*middle bar*) during mandatory breaths. *Upper bar*, neural Ti (gray area) and neural Te (white) for the group data are presented. *Middle bar*, periods describing ventilator timing are displayed, including trigger delay and ventilator Ti. *Bottom bar*, periods of infant-ventilator synchrony (white) and asynchrony (black).

# Excellent Reference on Edi Monitoring

Hindawi Publishing Corporation  
Critical Care Research and Practice  
Volume 2013, Article ID 384210, 7 pages  
<http://dx.doi.org/10.1155/2013/384210>

## *Review Article*

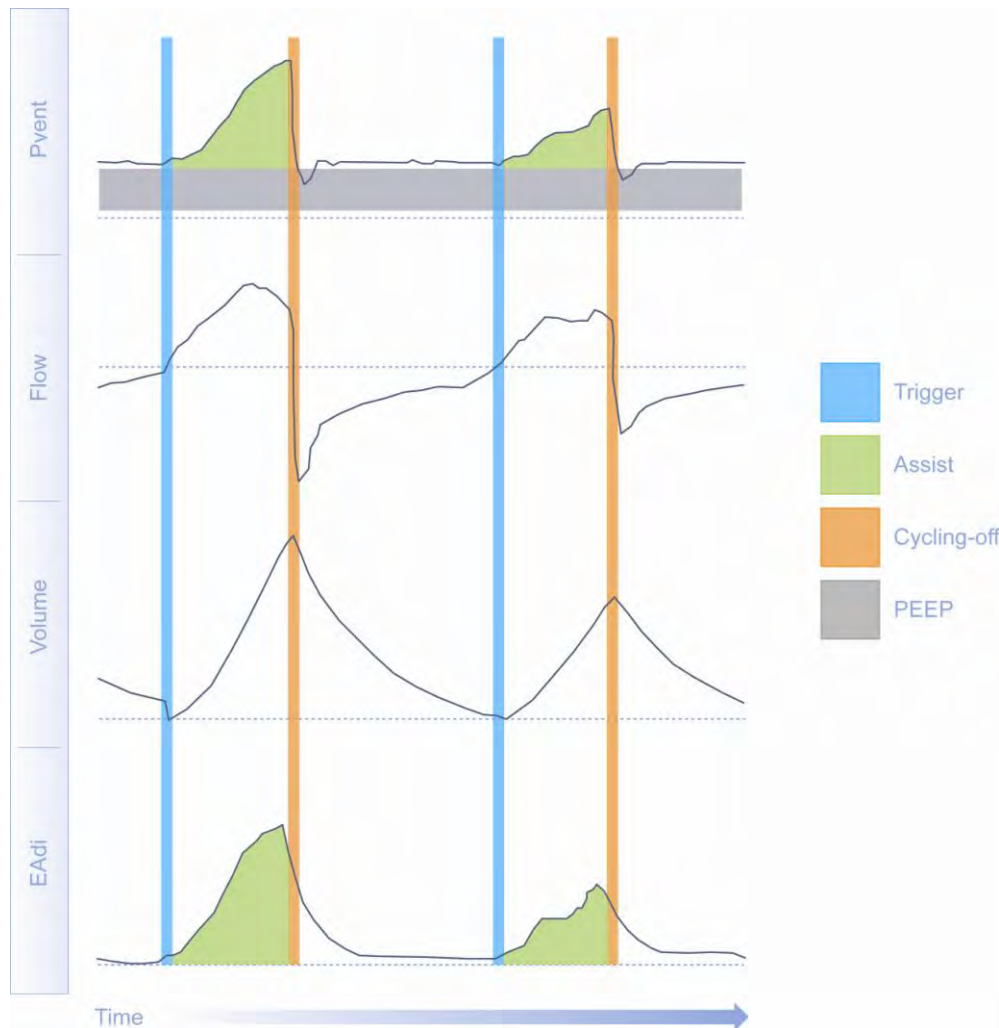
## **Interest of Monitoring Diaphragmatic Electrical Activity in the Pediatric Intensive Care Unit**

**Laurence Ducharme-Crevier, Geneviève Du Pont-Thibodeau, and Guillaume Emeriaud**

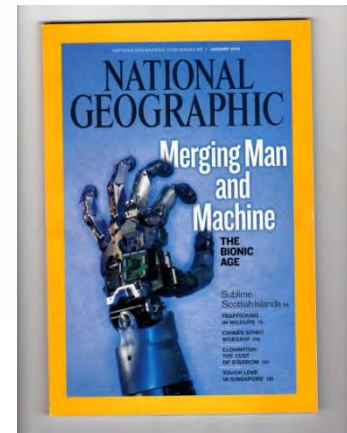
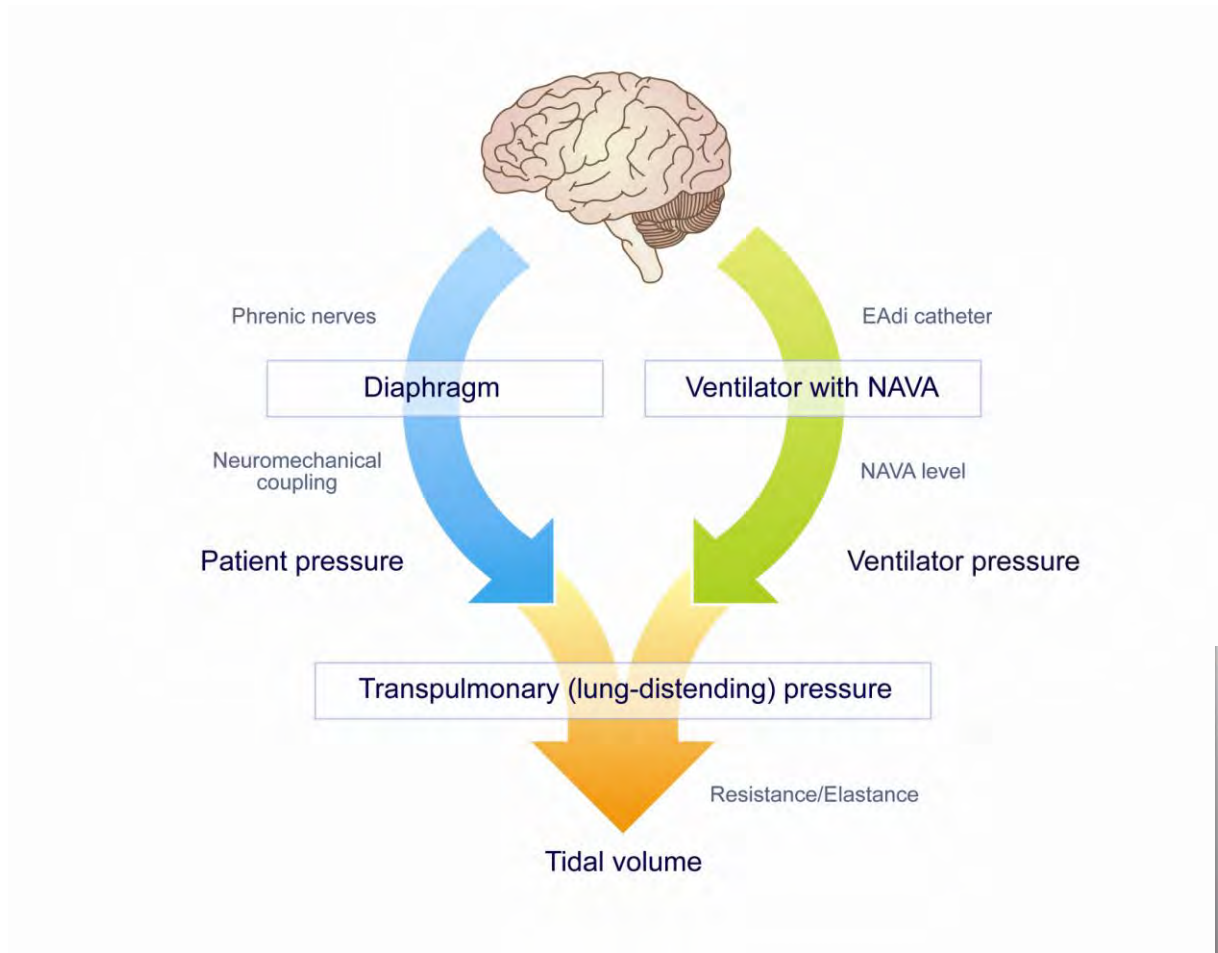
*Pediatric Intensive Care Unit, CHU Sainte-Justine, Université de Montréal, 3175 Chemin de la Côte Sainte-Catherine, Montreal, QC, Canada H3T 1C5*

Correspondence should be addressed to Guillaume Emeriaud; [guillaume.emeriaud@umontreal.ca](mailto:guillaume.emeriaud@umontreal.ca)

# Edi and Controlling Mechanical Ventilation: NAVA



# NAVA is an Artificial Respiratory Muscle





# Patient-Ventilator Interaction During Neurally Adjusted Ventilatory Assist in Low Birth Weight Infants

JENNIFER BECK, MAUREEN REILLY, GIACOMO GRASSELLI, LUCIA MIRABELLA, ARTHUR S. SLUTSKY, MICHAEL S. DUNN, AND CHRISTER SINDERBY (*Pediatr Res* 65: 663–668, 2009)

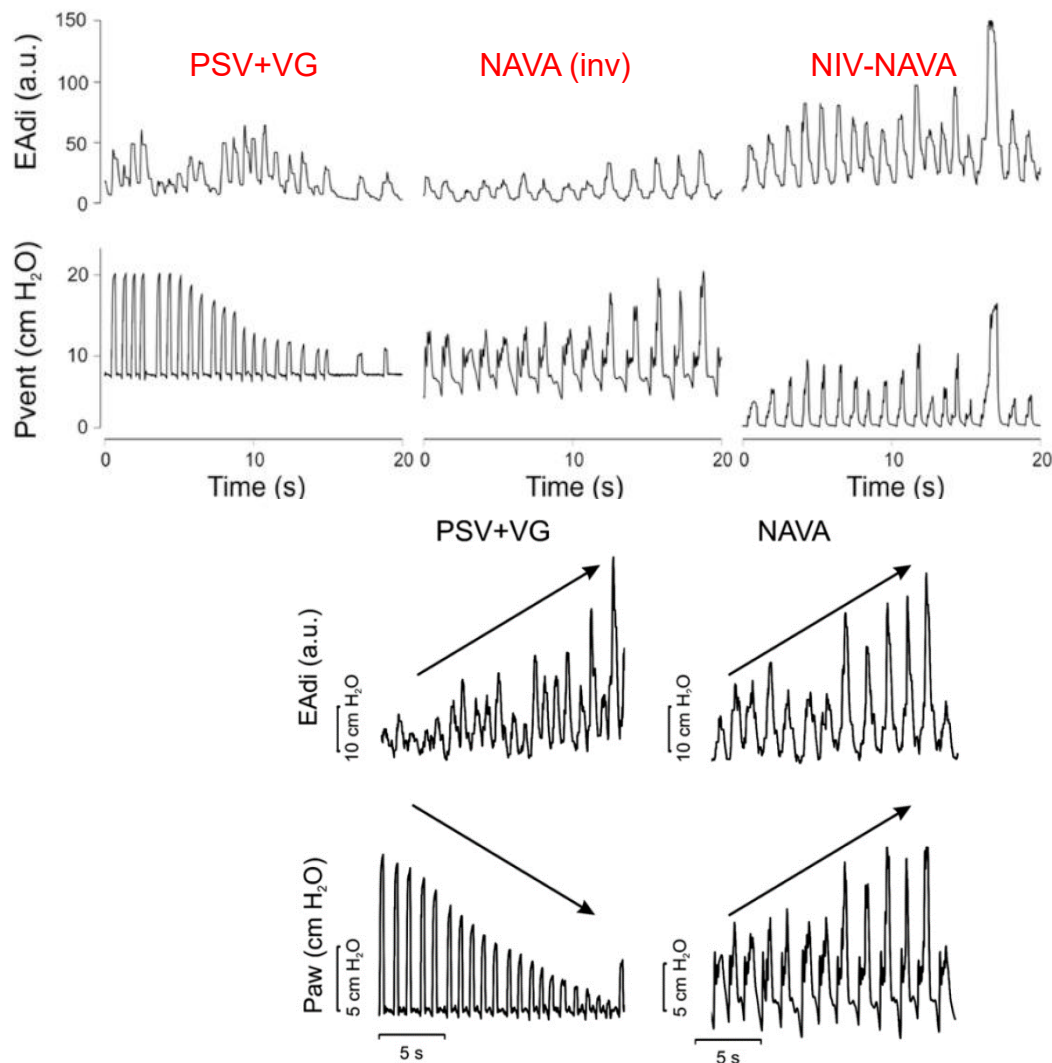


N = 7

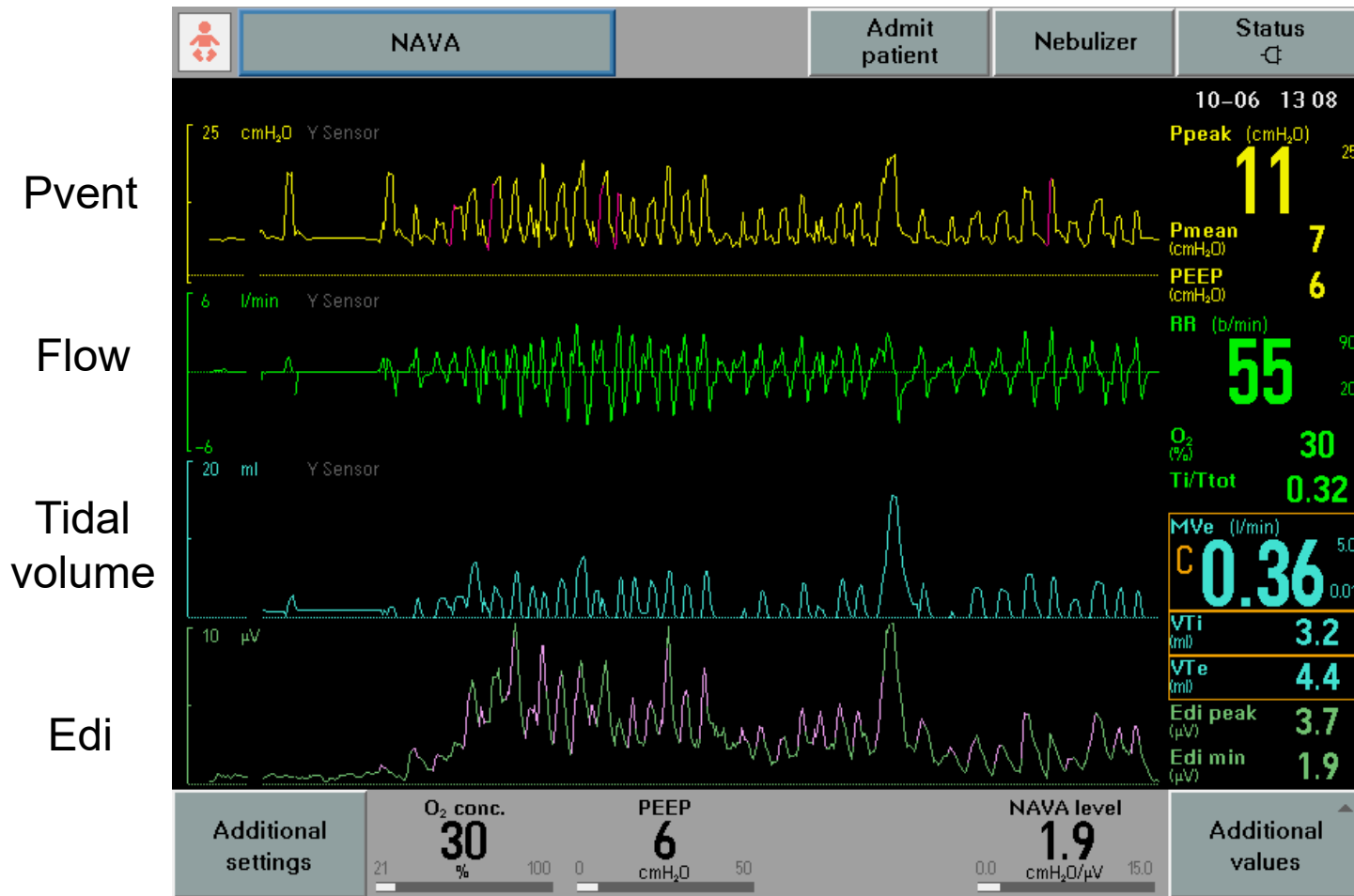
Mean weight = 976 g

Mean age = 12 days

GA at birth: 26 wks

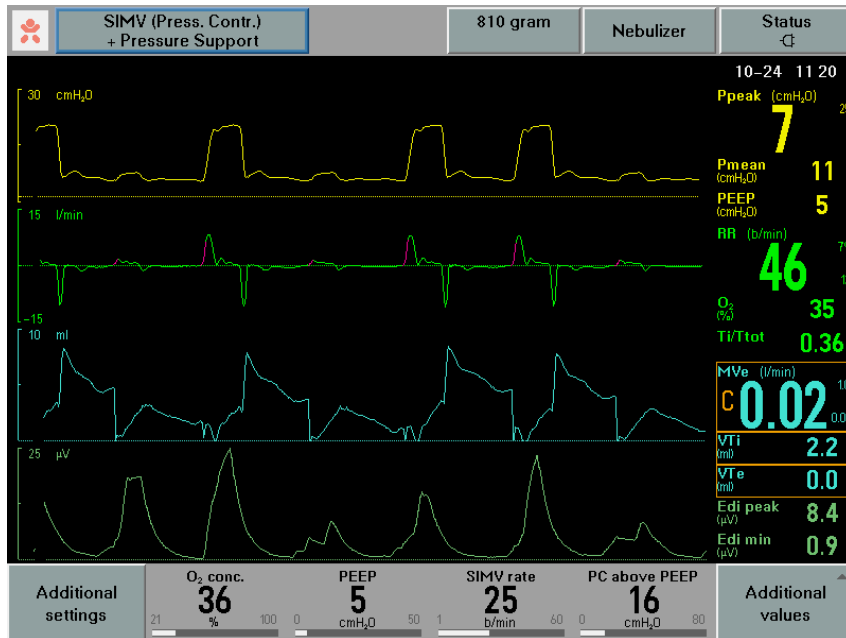


# NAVA in the preterm

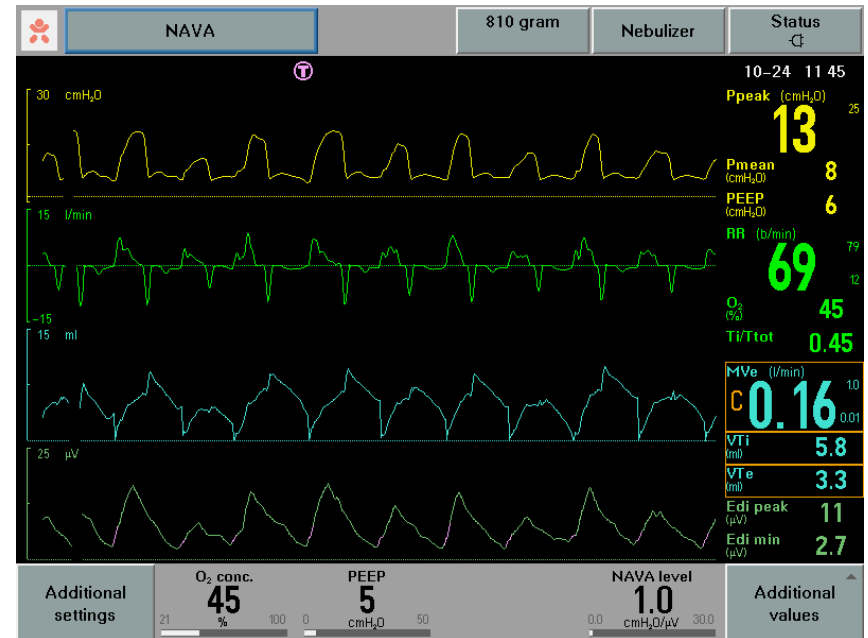


# NAVA improves synchrony (810 g)

SIMV



NAVA



# Neurally adjusted ventilatory assist improves patient–ventilator interaction in infants as compared with conventional ventilation

Alice Bordessoule<sup>1</sup>, Guillaume Emeriaud<sup>1</sup>, Sylvain Morneau<sup>1</sup>, Philippe Jouvett<sup>1</sup> and Jennifer Beck<sup>2,3</sup>

Pediatric RESEARCH

Volume 72 | Number 2 | August 2012

**Table 3.** Patient–ventilator interaction and respiratory variability

	NAVA	PCV	PSV	P value <sup>a</sup>
Patient–ventilator interaction				
Trigger delay (ms)	93 (20)	193 (87)	135 (29)	$P < 0.001$ – PCV vs. NAVA and PSV vs. NAVA
Cycling-off delay (ms)	17 (13)	12 (176)	–77 (81)	NS
→ Asynchrony index (%)	11 (3)	24 (11)	25 (9)	$P < 0.001$ – PCV vs. NAVA and PSV vs. NAVA
→ Wasted efforts (%)	0 (0)	4.3 (4.6)	6.5 (7.7)	$P < 0.05$ – PSV vs. NAVA
→ Percentage of breaths cycled off too early (%)	0.3 (0.4)	12 (13)	21 (19)	$P < 0.01$ – PCV vs. NAVA and PSV vs. NAVA
Correlation between peak Pvent and peak EAdi				
Determination coefficient R <sup>2</sup>	0.71 (0.22)	0.15 (0.16)	0.12 (0.12)	$P < 0.001$ – PCV vs. NAVA and PSV vs. NAVA
Slope	1.45 (1.5)	0.07 (0.1)	0.06 (0.04)	$P < 0.01$ – PCV vs. NAVA and PSV vs. NAVA
Respiratory variability				
→ Peak Edi – CV (%)	49 (27)	50 (29)	51 (32)	NS
Tidal volume – CV (%)	31 (26)	15 (12)	20 (15)	$P = 0.17$
→ Peak Pvent – CV (%)	24 (8)	2 (1)	2 (2)	$P < 0.01$ – PCV vs. NAVA and PSV vs. NAVA

Means (SD) are presented.

N = 10

Mean weight = 4.3 kg

Mean age = 2 mos

GA at birth: 26 wks

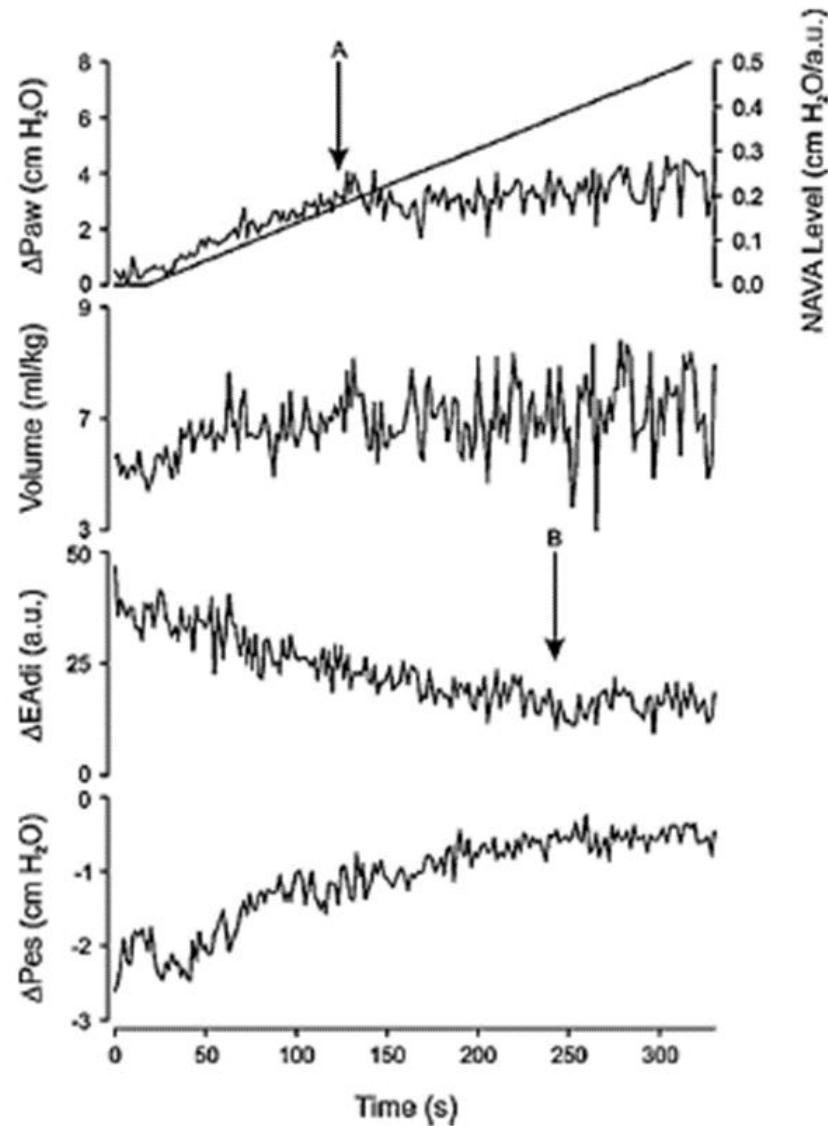
- 22 studies (PUBMED) in 305 patients of all ages have shown that NAVA improves synchrony compared to conventional ventilation modes
- 19 studies in 291 patients of all ages report improved or equivalent physiological parameters

# Impact of improved synchrony with NAVA (vs. Conv)

- Improved sleep (Delisle 2011)
- Improved COMFORT (less distress) (De Oliva 2012)
- Less central apnea (Mally 2013 PAS 2013; Delisle 2013)
- Tendency for less sedation (Allander 2013, ESPNIC 2013)
- Equivalent or improved blood gases ( $n > 5$  for improvement, rest equivalent when measured)
- More efficient ventilation (less P required) ( $n > 13$  studies)
- Less work of breathing to trigger (Spahija 2010, Clement 2011)
- Improved variability of delivered assist and  $V_t$  ( $n \geq 3$ )
- Improved lung aeration (Blankman 2013)



# NAVA Can Prevent Over-Assist



# Titration and Implementation of Neurally Adjusted Ventilatory Assist in Critically Ill Patients\*

Lukas Brander, MD; Howard Leong-Poi, MD; Jennifer Beck, PhD;  
Fabrice Brunet, MD; Stuart J. Hutchison, MD; Arthur S. Slutsky, MD;  
and Christer Sinderby, PhD

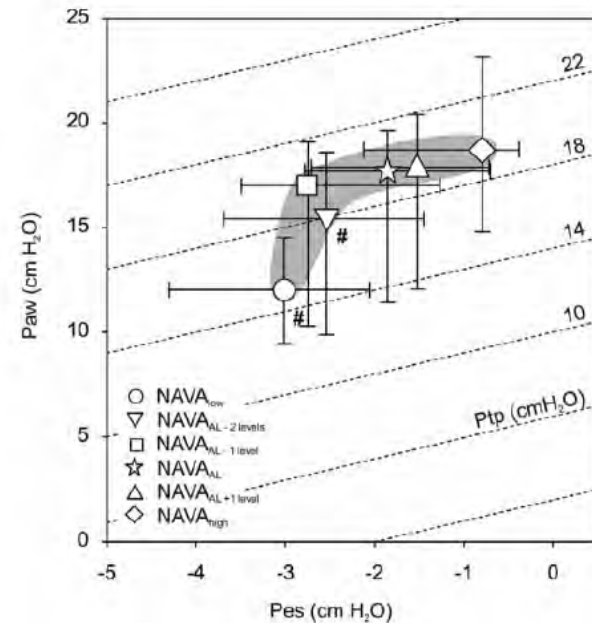
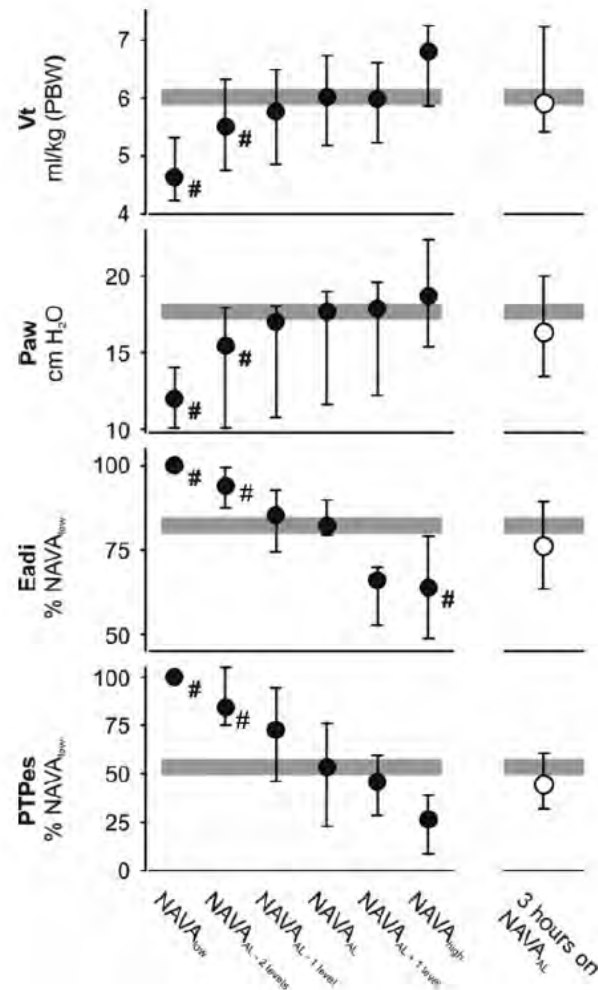
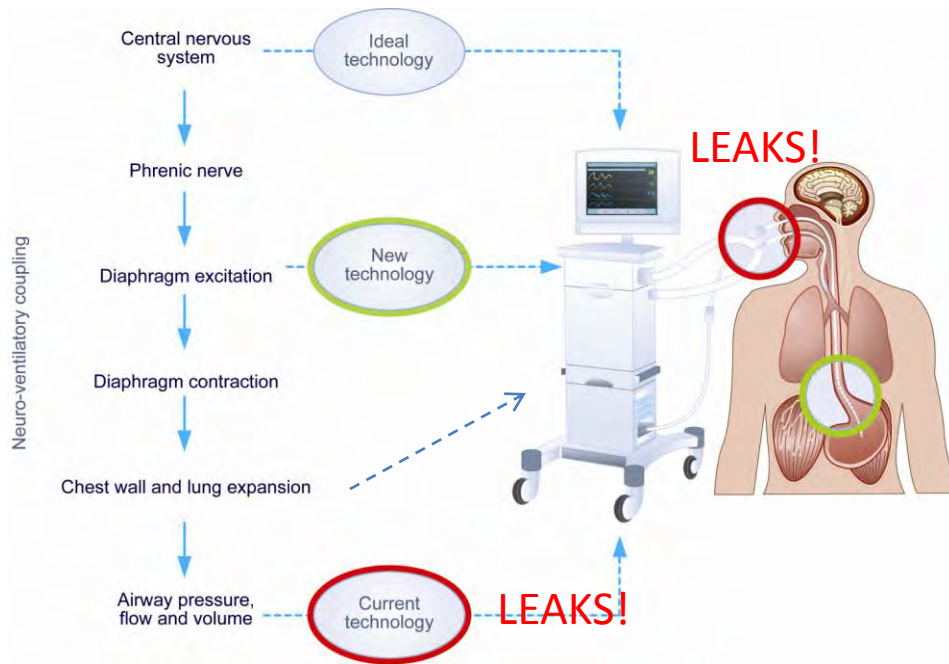


FIGURE 3. Group data for the interaction among mean inspiratory  $P_{aw}$  including PEEP,  $P_{es}$ , and  $P_{tp}$  during the NAVA level titration. From NAVA<sub>low</sub> to one level below NAVA<sub>AL</sub> (NAVA<sub>AL</sub>-1 level),  $P_{aw}$  increased by 5.0 cm H<sub>2</sub>O and reduced the  $P_{es}$  deflection by 0.5 cm H<sub>2</sub>O such that  $P_{tp}$  increased by 4.5 cm H<sub>2</sub>O. Further increasing the NAVA level from NAVA<sub>AL</sub> to NAVA<sub>high</sub> resulted in changes of  $P_{aw}$  and  $P_{es}$  that were similar in magnitude ( $P_{aw}$  increased by 1.7 cm H<sub>2</sub>O and  $P_{es}$  decreased by 1.95 cm H<sub>2</sub>O, respectively) and hence in an essentially unaltered  $P_{tp}$ . The PEEP was 8.5 cm H<sub>2</sub>O (quartiles, 5.3 to 11.8 cm H<sub>2</sub>O) and remained unchanged during the NAVA level titration. Abbreviations are the same as for Figure 2. Symbols represent group median, lines indicate the 25th and 75th percentiles. # =  $p < 0.05$  vs NAVA<sub>AL</sub>.

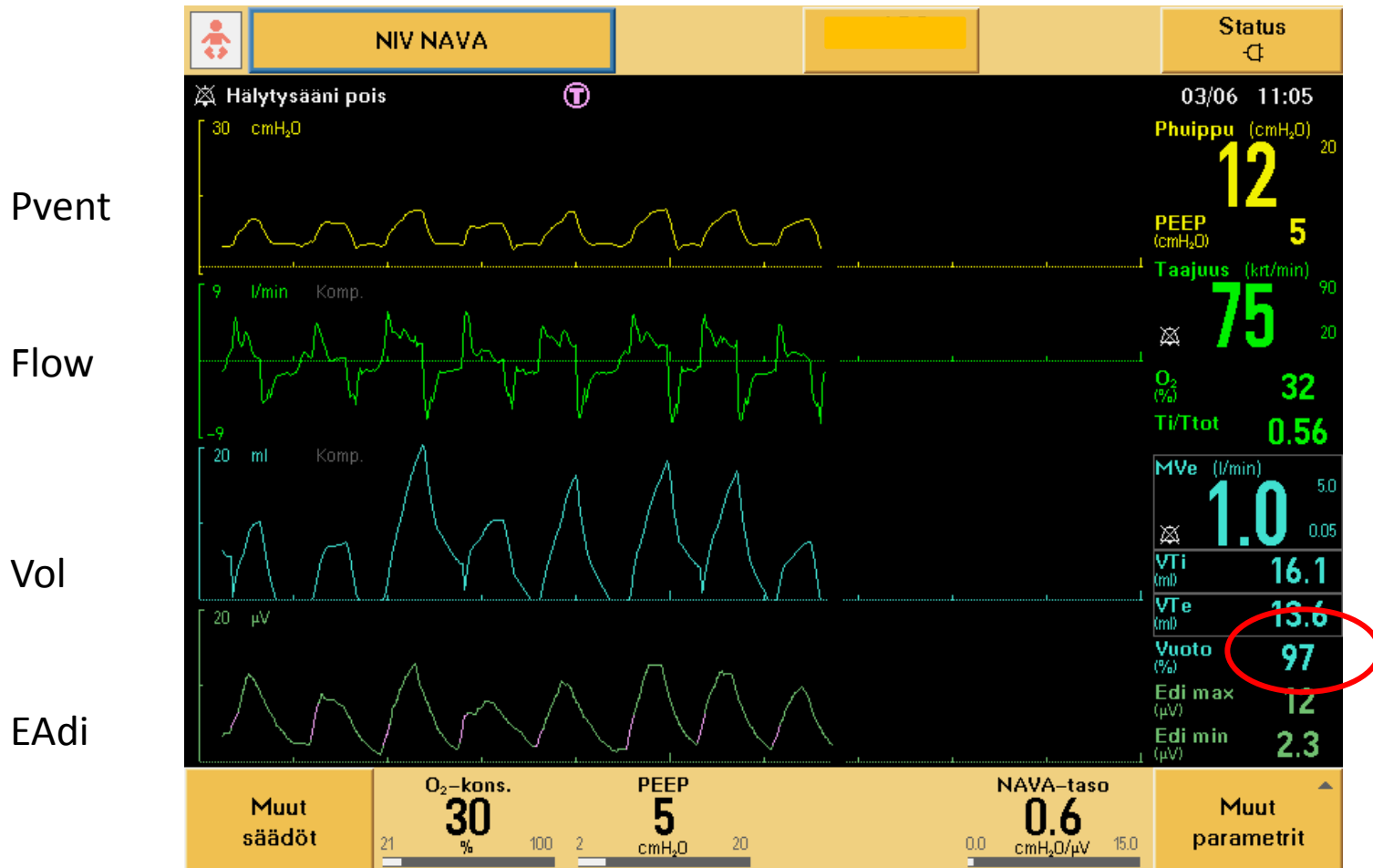
- 7 studies in 100 patients show down-regulation of Edi with increasing NAVA
- 14 studies in 201 adults on NAVA show mean Vt 6.9 ml/kg, rr 24 bpm
- 13 studies in 197 infants on NAVA show mean Vt 6.3 ml/kg, rr 45 bpm

# Factors Affecting NIV



- Controller signal for patient-ventilator synchrony
- Inadequate (pneumatic) monitoring
- Properties of interface
  - Masks and prongs (leaks)
  - Prongs (resistance)
- Leaks and pressure delivery
- Upper airways (protection and control of FRC)

# NIV-NAVA in the preterm



## Patient-Ventilator Interaction During Neurally Adjusted Ventilatory Assist in Low Birth Weight Infants

JENNIFER BECK, MAUREEN REILLY, GIACOMO GRASSELLI, LUCIA MIRABELLA, ARTHUR S. SLUTSKY,  
 MICHAEL S. DUNN, AND CHRISTER SINDERBY

**Table 2.** Vital signs, ventilator parameters, neural breathing pattern, and patient-ventilator interaction

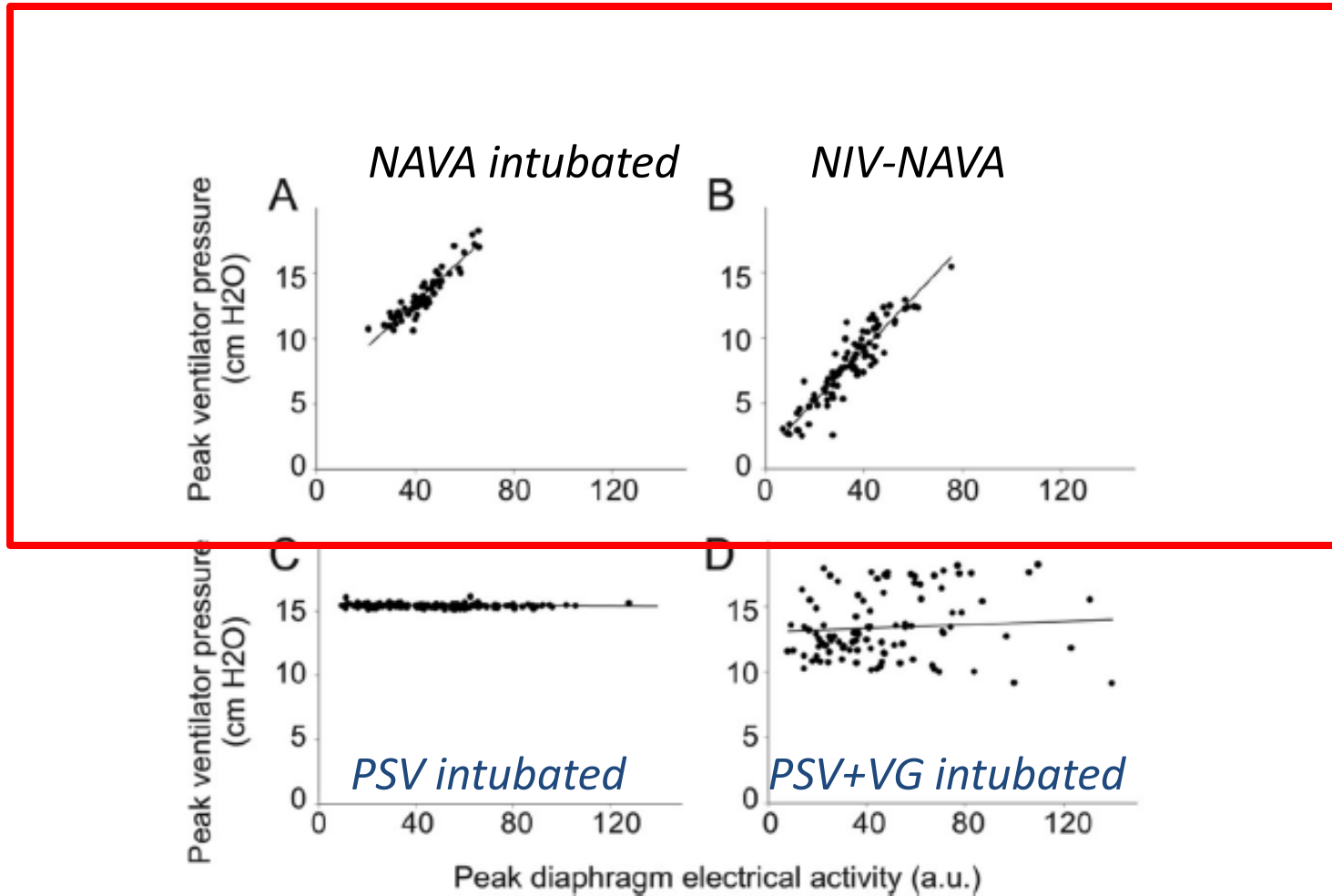
	Conventional ventilation	NAVA-intubated	NAVA-extubated	<i>p</i>
	<i>n</i> = 7	<i>n</i> = 7	<i>n</i> = 5	
FIO <sub>2</sub> (%)	25.1 (5.8)†	26.4 (5.9)*	38.0 (19.1)	0.0034
SAO <sub>2</sub> (%)	94.6 (2.7)	94.1 (3.5)	95.4 (6.5)	0.961
Tco <sub>2</sub> (mm hg)	53.4 (14.6)	52.6 (13.0)	60.6 (12.5)	0.787
Heart rate (per min)	160 (15)	159 (11)	166 (11)	0.868
	<i>n</i> = 5	<i>n</i> = 7	<i>n</i> = 5	
MAPi (cm H <sub>2</sub> O)	12.5 (1.5)*	9.6 (1.8)*	5.5 (1.6)	0.002
ΔPi (cm H <sub>2</sub> O)	9.3 (1.3)	9.9 (1.3)	9.4 (3.1)	0.710
EAdi phasic (au)	27.9 (19.5)	43.6 (18.7)	44.8 (32.4)	0.333
EAdi tonic (au)	5.5 (2.0)	4.7 (3.0)	4.7 (1.4)	0.157
Nti (msec)	258 (43)	406 (131)	436 (198)	0.09
Nte (msec)	712 (139)	875 (237)†	1001 (256)	0.044
Nrr (per min)	74 (7)	54 (14)†	51 (14)†	0.004
EAdi-time product (au*s/min)	556.4 (421.2)	823.4 (444.3)	670.2 (524.5)	0.504
Trigger delay (ms)	74 (17)	72 (23)	76 (33)	0.698
Cycling-off delay (ms)	-120 (66)	32 (34)†	28 (11)†	<0.001
R <sup>2</sup> for EAdi vs Pvent	0.08 (0.1)	0.80 (0.06)†	0.73 (22)†	<0.001
Slope EAdi vs Pvent (cm H <sub>2</sub> O per au)	0 (0.01)	0.19 (0.1)†	0.2 (0.1)†	0.007

\* Statistically different from NAVA-ext.

† Statistically different from Conv.

FIO<sub>2</sub>, fraction of inspired oxygen; SAO<sub>2</sub>, oxygen saturation; Tco<sub>2</sub>, transcutaneous carbon dioxide; MAPi, mean inspiratory airway pressure; ΔPi, delta inspiratory pressure above PEEP; EAdi, electrical activity of the diaphragm; Nti, neural inspiratory time; Nte, neural expiratory time; Nrr, neural respiratory rate; R<sup>2</sup>, determination coefficient; Pvent, ventilator delivered pressure.

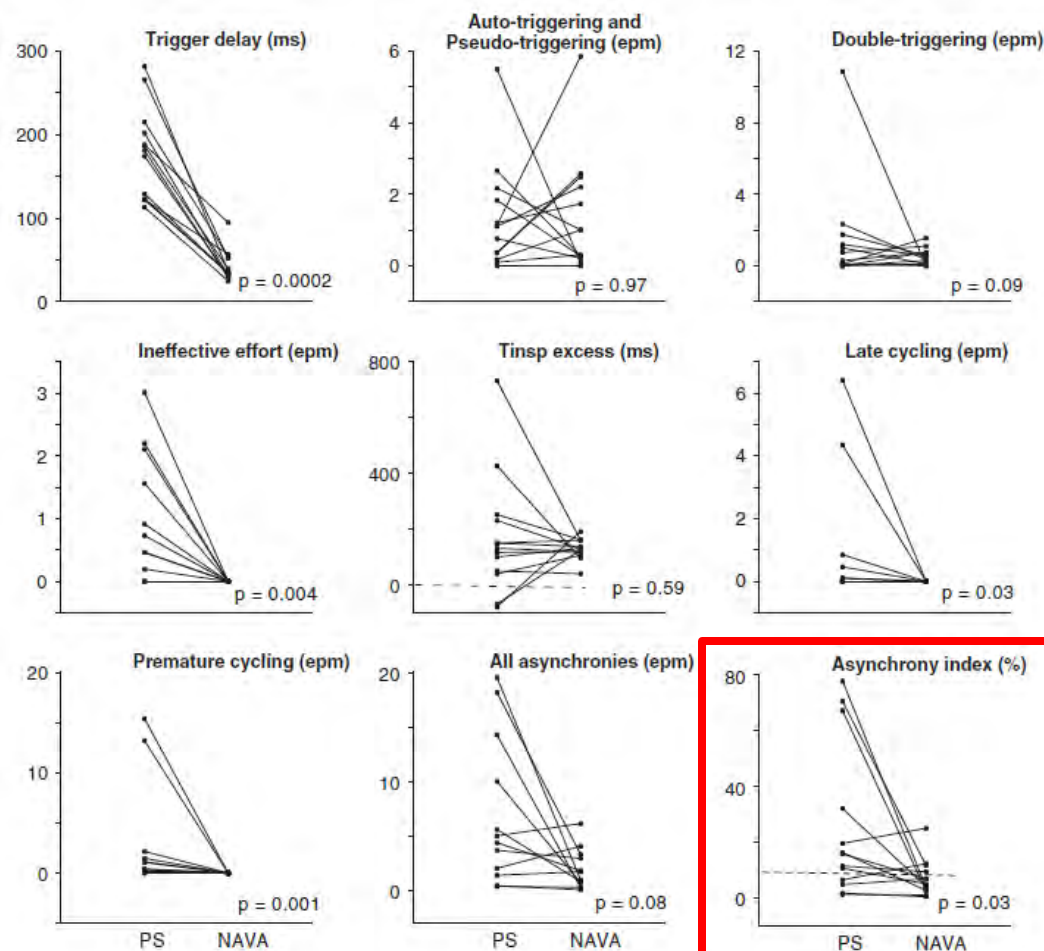
# Proportionality between patient effort and ventilatory assist





Lise Piquilloud  
Didier Tassaux  
Emilie Bialais  
Bernard Lambermont  
Thierry Sottiaux  
Jean Roeseler  
Pierre-François Laterre  
Philippe Joliet  
Jean-Pierre Revelly

# **Neurally adjusted ventilatory assist (NAVA) improves patient-ventilator interaction during non-invasive ventilation delivered by face mask**

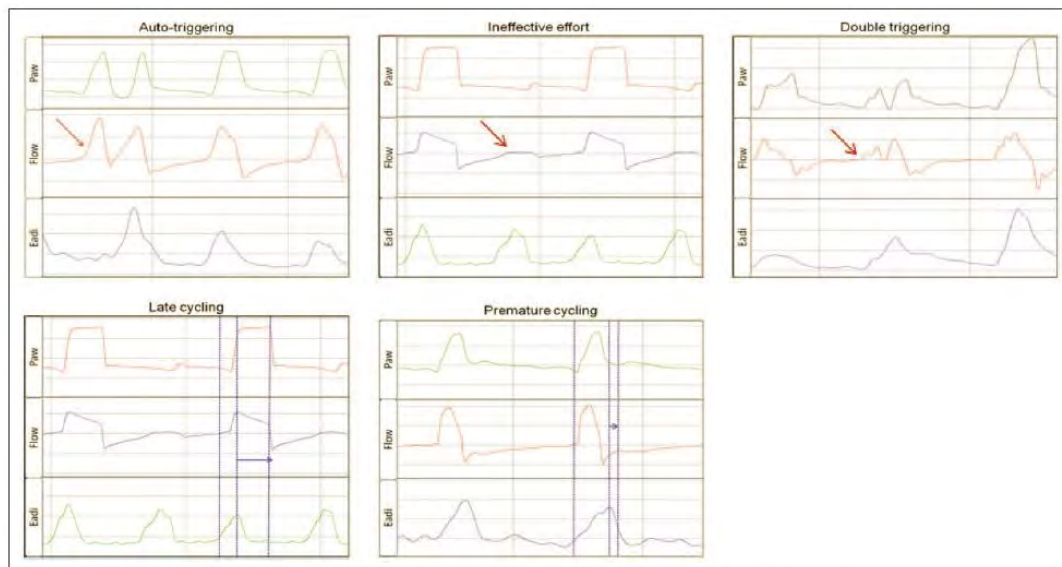


N= 13 Adult ICU (incl 2 COPD)  
20 min NIVNAVA, 20 MIN NIV PSV  
Mask

# Patient–Ventilator Asynchrony During Noninvasive Pressure Support Ventilation and Neurally Adjusted Ventilatory Assist in Infants and Children

PCCM, In Press, June 2013

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**Figure 2.** Representative tracings of the five types of asynchrony. Eadi = electrical activity of the diaphragm tracing, Paw= airway pressure, flow = instantaneous flow. Downward pointing arrows indicate relevant events.

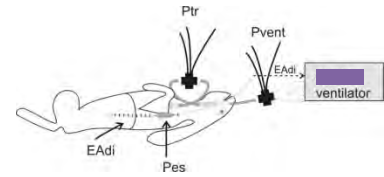
N=6  
Consecutive, requiring NIV for ARF  
Age: 18 mos  
Weight: 8 kg  
Mask (4) or prongs (2)

**TABLE 2. Number and Type of Asynchronies (n = 6)**

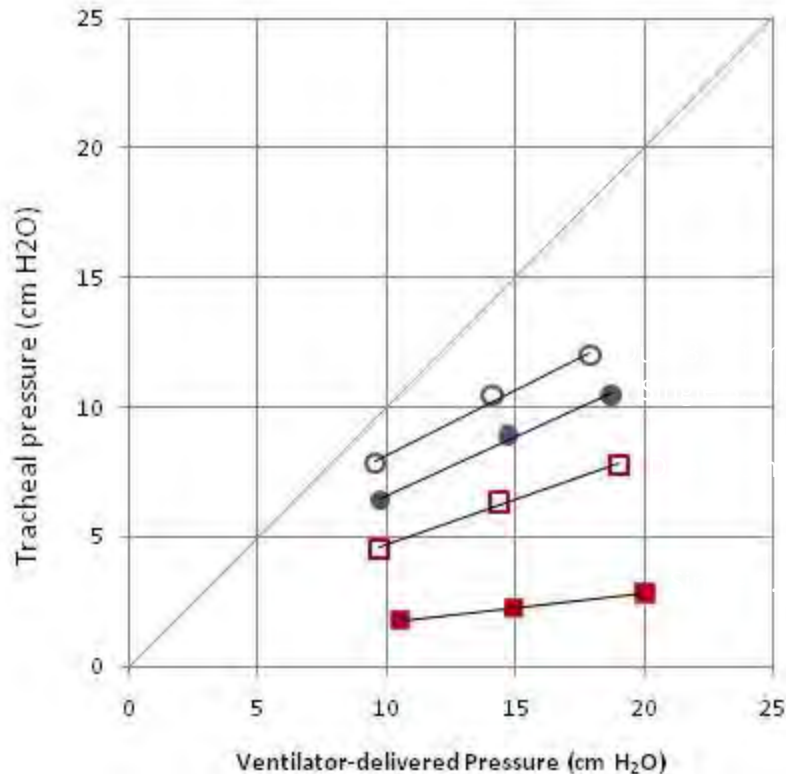
	PSinit	PSbest	NAVA	<i>p</i> Analysis of Variance	<i>p</i> NAVA vs PSinit	<i>p</i> NAVA vs PSbest	<i>p</i> PSbest vs PSinit
Autotriggering (n/min)	4.8 (1.7–12)	4.7 (1.3–7)	0.2 (0–0.9)	0.083	—	—	—
Ineffective efforts (n/min)	9.9 (1.7–18)	5.1 (1–15.6)	0 (0–0)	0.029	<0.05	NS	<0.05
Double triggering (n/min)	0.2 (0.1–0.3)	0 (0–0.8)	0.7 (0.2–1.7)	0.120	—	—	—
Late cycling (n/min)	0.5 (0.1–1.7)	0 (0–0.8)	0 (0–0.2)	0.430	—	—	—
Premature cycling (n/min)	6.3 (3.2–18.7)	3.4 (1.1–7.7)	0 (0–0)	0.022	<0.05	NS	NS
Asynchrony index (%)	65.5 (42–76)	40 (28–65)	2.3 (0.7–5)	<0.001	<0.05	<0.05	NS

PS = pressure support, PSinit = mode in pressure support with initial adjustment of the cycling criterion, PSbest = mode in pressure support with the cycling criterion that allowed the best results of asynchrony, NAVA = neurally adjusted ventilatory assist.  
Values are expressed in median (25<sup>th</sup>–75<sup>th</sup> percentile).

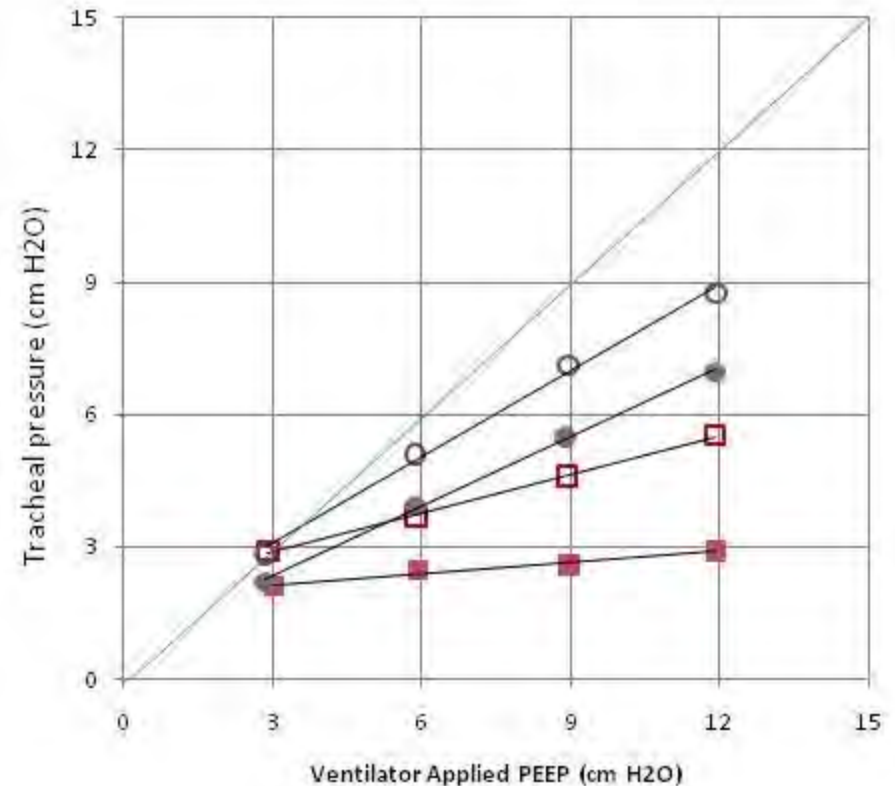
# Effect of nasal prongs on pressure delivery (n=10)



NAVA Titration



PEEP Titration

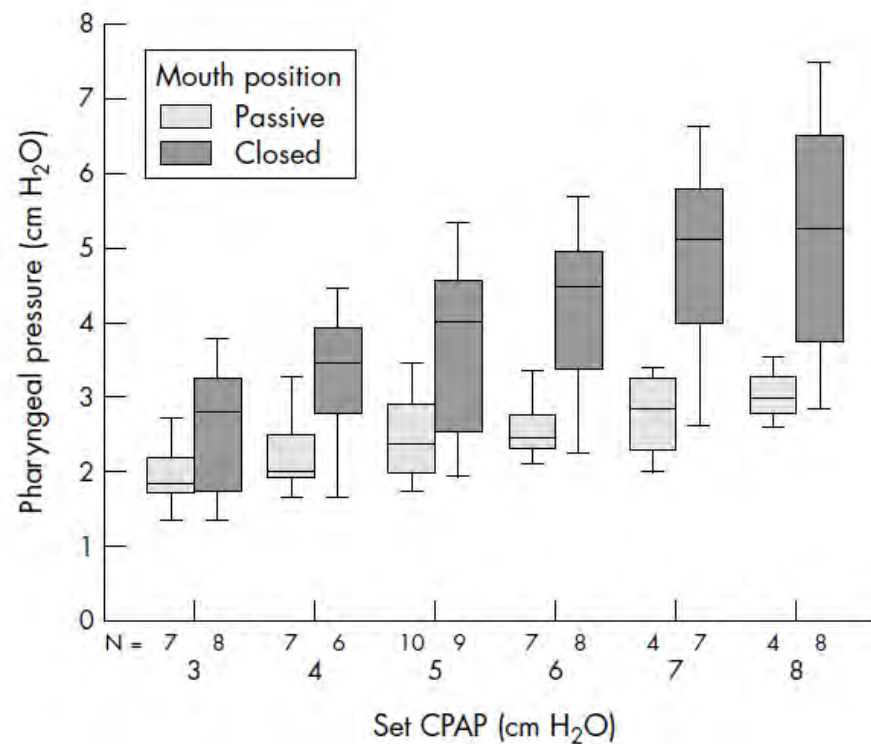


## SHORT REPORT

# Pharyngeal pressure in preterm infants receiving nasal continuous positive airway pressure

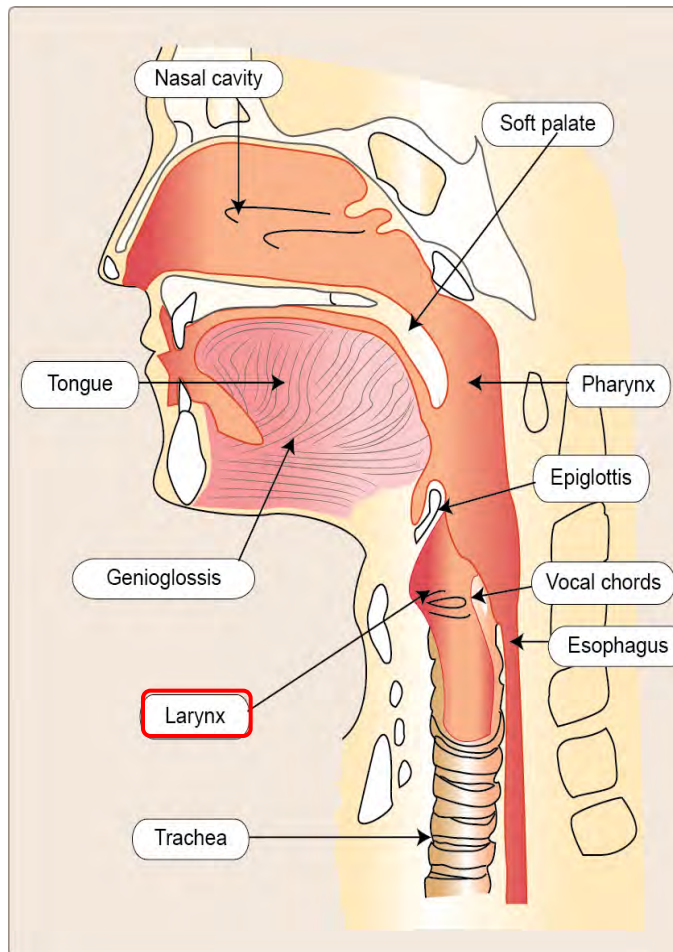
A G De Paoli, R Lau, P G Davis, C J Morley

*Arch Dis Child Fetal Neonatal Ed* 2005;**90**:F79-F81. doi: 10.1136/adc.2004.052274



N= 11 preterms  
Binasal Hudson prongs  
Bubble cpap

# Upper Airways

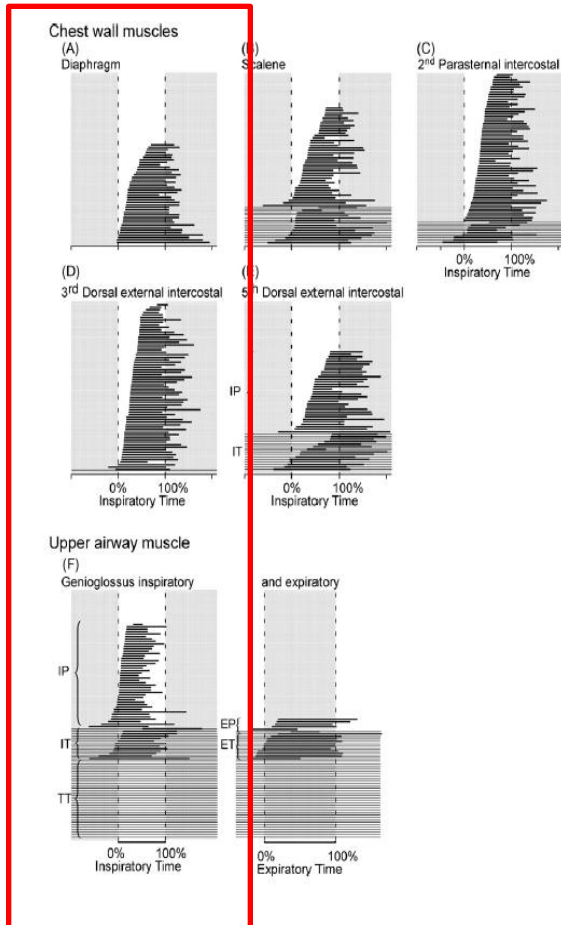


## Role:

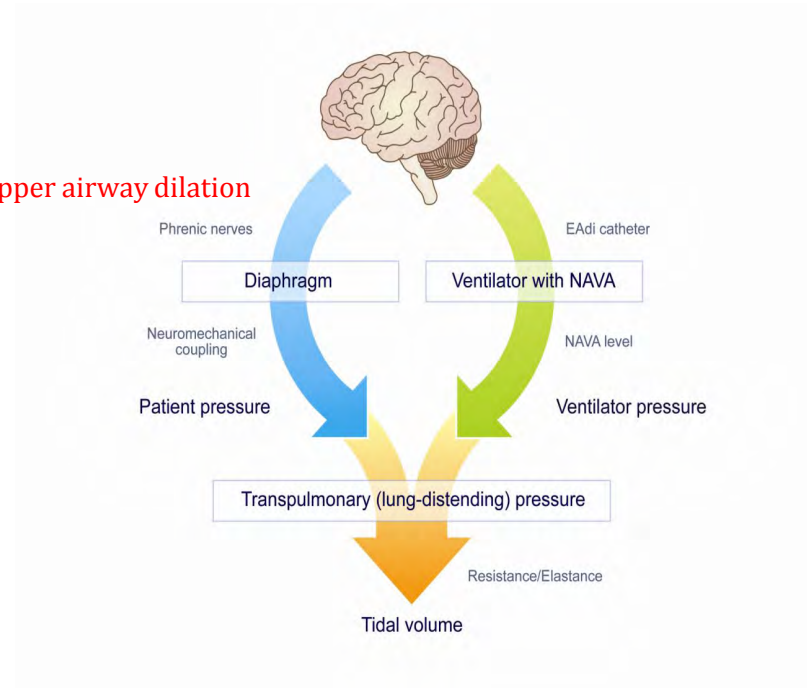
- Humidification
- Speech
- Swallowing
- Airway protection
- **Airway dilation for inspiration**
- Braking of expiratory flow to maintain EELV



# Timing of Activation of Upper Airways

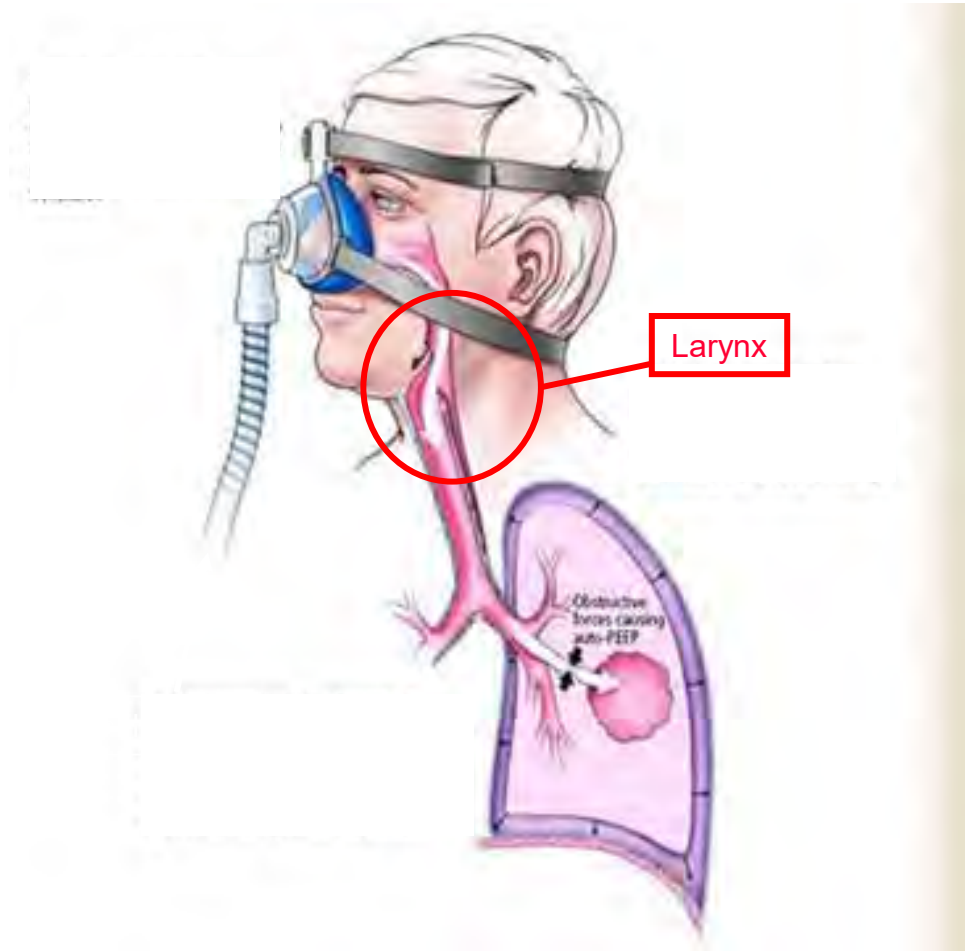


## Upper airway dilation



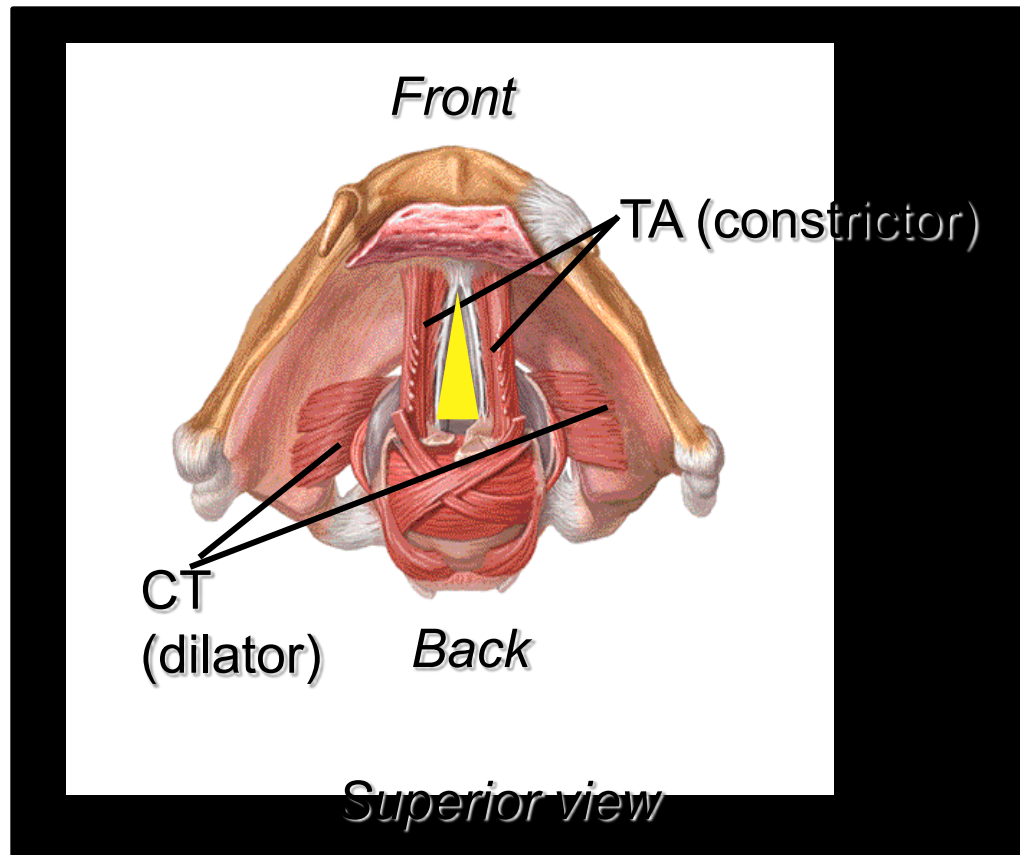
# Larynx and NIV

- Larynx = a closing valve
- Original function = to defend the lower airways against potentially harmful intruders



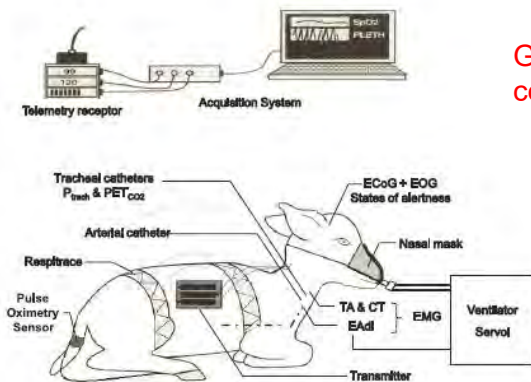


# Important Laryngeal Muscles



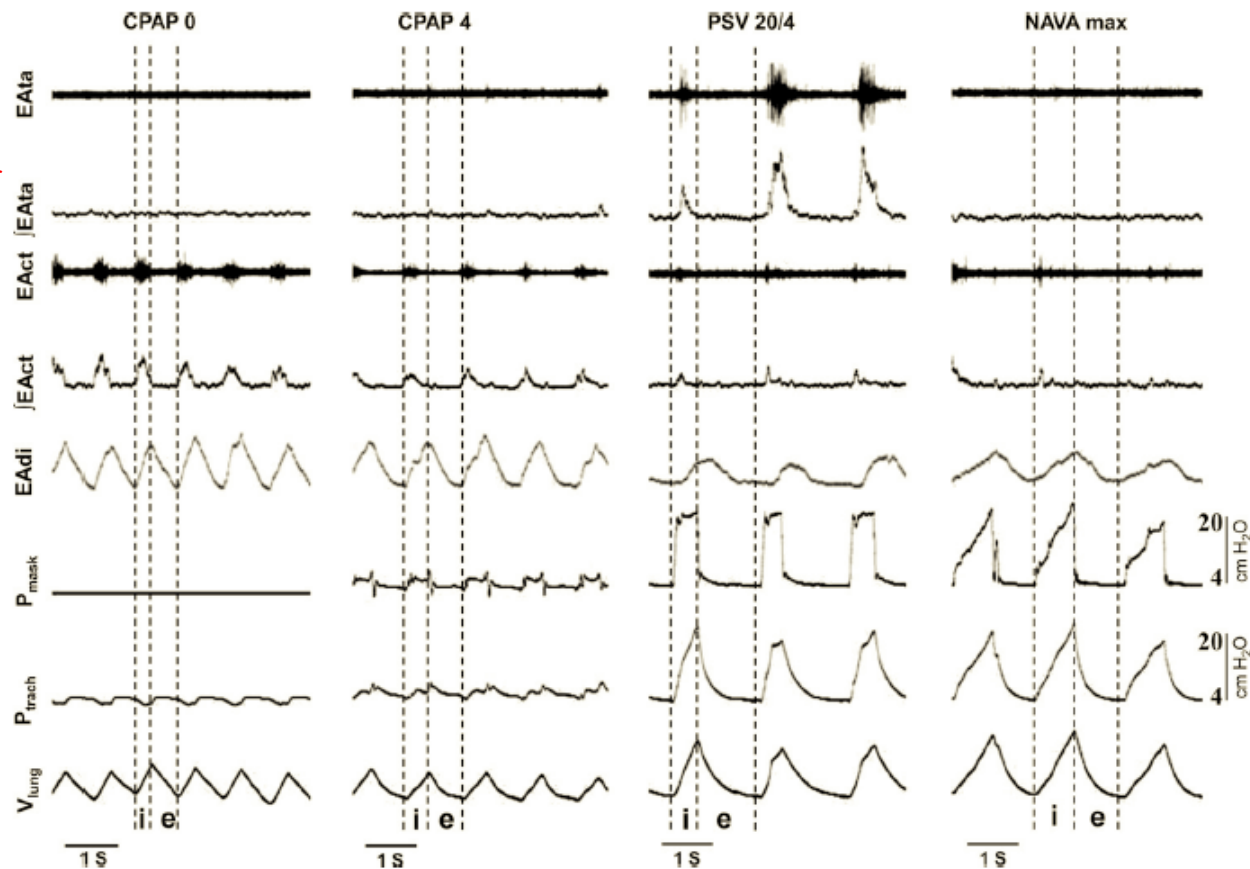
# Absence of inspiratory laryngeal constrictor muscle activity during nasal neurally adjusted ventilatory assist in newborn lambs

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Glottal  
constrictor

Glottal  
dilator



# Summary

- The Edi is a physiological signal representative of central respiratory output
- The Edi is normally present in spontaneously breathing subjects. The waveform has a characteristic cyclic/phasic pattern with quantifiable measures of amplitude and timing.
- The Edi is essentially a vital sign, just like the electrocardiogram.
- With Edi monitoring, clinicians can answer these questions:
  - Is my patient breathing?
  - Is my patient synchronous with the ventilator?
  - Is my treatment/intervention helping?
  - Is it a central or obstructive apnea?
- NAVA provides synchronized and proportional assist (invasive and non-invasive)
- Ventilator becomes a second respiratory muscle and is therefore a slave to the patient's own protective reflexes and control of breathing responses